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- Key terms
              FICS' ENTERED AT 15:27:39 ON 06 MAR 2002)
           2908 SEA FILE=CAPLUS ABB=ON PLU=ON (DETERM? OR DETECT? OR
L1
               DET## OR SCREEN?) (5A) PATHOGEN##
            105 SEA FILE=CAPLUS ABB=ON PLU=ON L1 AND SUBSTRATE
L2
              7 SEA FILE=CAPLUS ABB=ON PLU=ON L2 AND CRYSTAL?
           2908 SEA FILE=CAPLUS ABB=ON PLU=ON
                                               (DETERM? OR DETECT? OR
L1
               DET## OR SCREEN?) (5A) PATHOGEN##
           1380 SEA FILE=CAPLUS ABB=ON PLU=ON L1 AND (METHOD OR
L6
               TECHNIQUE OR APPARAT? OR DEVICE)
             60 SEA FILE=CAPLUS ABB=ON PLU=ON L6 AND SUBSTRATE
            15 SEA FILE=CAPLUS ABB=ON PLU=ON L7 AND BIND?
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L9 ANSWER 1 OF 20 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:107584 CAPLUS

DOCUMENT NUMBER: 136:131210

TITLE: A device for detecting bacterial contamination

and method of use

INVENTOR(S): Sanders, Mitchell C.

PATENT ASSIGNEE(S): Expressive Constructs, Inc., USA

SOURCE: PCT Int. Appl., 25 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATI	ENT 1	NO.		KI	ND	DATE			A.	PPLI	CATI	N NC	0.	DATE		
									_							
WO 2	2002	0104	33	A:	2	20020	0207		W	200	01-U	S146	13	2001	0503	
	W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,
														KR,		
														MX,		
		NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТĴ,	TM,	TR,	TT,
		TZ,	UA,	UG,	UZ,	VN,	YU,	ZA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,
		ТJ,	TM													
	RW:													ΑT,		
		CY,	DE,	DK,	ES,	FΙ,	FR,	GB,	GR,	ΙE,	ΙT,	LU,	MC,	NL,	PT,	SE,
		TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,
		ΤG														
DTTV	A DD:	דא ז	TNEO					1	FIG 21	000 <u>-</u> 1	2014	ሰፍ	D	2000	በፍሰገ	

PRIORITY APPLN. INFO.: US 2000-201405 P 20000503

A device and method for detecting the presence or absence of a prokaryotic microorganism are provided, comprising the steps of identifying a protein, such as a microbial-specific protease that characterizes the presence of a specific prokaryotic microbe and thereby provides a marker for that microbe; detecting the protease that is a marker for the presence of a specific prokaryotic microbe by cleaving a substance when the protease is present; and signaling the presence of that protease when cleavage has occurred. More specifically, the method comprises identifying at least one outer membrane protein or a secreted protein that is unique to a particular microbial pathogen such as for example Listeria monocytogenes and that is substrate specific.

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ANSWER 2 OF 20 CAPLUS COPYRIGHT 2002 ACS
L9
                              2001:816983 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                              135:354956
                              High density protein arrays for screening of
TITLE:
                              protein activity
                              Snyder, Michael; Reed, Mark; Zhu, Heng; Klemic,
INVENTOR(S):
                              James Frank
                              Yale University, USA
PATENT ASSIGNEE(S):
                              PCT Int. Appl., 69 pp.
SOURCE:
                              CODEN: PIXXD2
                              Patent
DOCUMENT TYPE:
                              English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                          KIND DATE
                                                    APPLICATION NO.
      PATENT NO.
                                                    _____
      _____
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                                              WO 2001-US14526 20010504
                         A1 20011108
      WO 2001083827
               AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TT, TM
                RU, TJ, TM
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD,
                TG
                                                                     P 20000504
PRIORITY APPLN. INFO.:
                                                US 2000-201921
                                                US 2000-221034 P 20000727
      The invention concerns protein chips useful for the large-scale
AB
      study of protein function where the chip contains densely packed
      reaction wells. The invention also relates to methods of
      using protein chips to assay simultaneously the presence, amt.,
      and/or function of proteins present in a protein sample or on one
      protein chip, or to assay the presence, relative specificity, and
      binding affinity of each probe in a mixt. of probes for each
      of the proteins on the chip. The invention also relates to
      methods of using the protein chips for high d. and small
      vol. chem. reactions. Also, the invention relates to polymers
      useful as protein chip substrates and methods of
      making protein chips. The invention further relates to compds.
      useful for the derivatization of protein chip substrates.
                                      THERE ARE 5 CITED REFERENCES AVAILABLE FOR
REFERENCE COUNT:
                              5
                                      THIS RECORD. ALL CITATIONS AVAILABLE IN
                                      THE RE FORMAT
     ANSWER 3 OF 20 CAPLUS COPYRIGHT 2002 ACS
L9
                               2001:614223 CAPLUS
ACCESSION NUMBER:
                              Method and apparatus for
TITLE:
                              detection of microscopic
                              pathogens
                              Abbott, Nicholas L.; Skaife, Justin J.
INVENTOR(S):
                              Wisconsin Alumni Research Foundation, USA
PATENT ASSIGNEE(S):
SOURCE:
                               PCT Int. Appl.
                              CODEN: PIXXD2
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Patent
DOCUMENT TYPE:
                          English
LANGUAGE:
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FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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APPLICATION NO.
                              DATE
                        KIND
     PATENT NO.
                              _____
                                              WO 2001-US4858
                              20010823
                                                                 20010215
                         A2
     WO 2001061357
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
              CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK,
              LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ,
              PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ,
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
              TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD,
                                           US 2000-182941
                                                              P 20000216
PRIORITY APPLN. INFO.:
     Detection apparatus for use in the detection of the
     presence of a selected pathogen in a sample are disclosed. Such
     apparatus includes: a substrate with a detection
     region on a surface thereof, the detection region having
     microstructures including grooves formed therein that will align
     liquid crystal material in contact therewith, the width
     and depth of the grooves being in the range of 10 .mu.m or less; a
     blocking layer on the surface of the detection region of the
     substrate that does not disrupt the alignment of liquid
     crystal material in contact therewith, the blocking layer
     blocking nonspecific adsorption of pathogens to the surface; and a
     binding agent on the surface of the detection region of the
     substrate, the binding agent specifically
     binding the selected pathogen.
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ANSWER 4 OF 20 CAPLUS COPYRIGHT 2002 ACS
L9
                        2000:792858 CAPLUS
ACCESSION NUMBER:
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134:69953 DOCUMENT NUMBER:

Micromosaic Immunoassays TITLE:

Bernard, Andre; Michel, Bruno; Delamarche, AUTHOR(S):

Emmanuel

Zurich Research Laboratory, IBM Research, CORPORATE SOURCE:

Rueschlikon, CH-8803, Switz.

Analytical Chemistry (2001), 73(1), 8-12 CODEN: ANCHAM; ISSN: 0003-2700 SOURCE:

American Chemical Society PUBLISHER:

DOCUMENT TYPE: Journal English LANGUAGE:

Immunoassays are widely used for medical diagnostics and constitute the principal method of detecting pathogenic agents and thus of diagnosing many diseases. These assays, which are most often performed in well plates, would be greatly improved by a practical method to pattern a series of antigens on a flat surface and to localize their binding to many analytes. But no obvious method exists to expose a planar surface successively to a series of antigens and analytes. Here, we present miniaturized mosaic immunoassays based on patterning lines of antigens onto a surface by

means of a microfluidic network (.mu.FN). Solns. to be analyzed are delivered by the channels of a second .mu.FN across the pattern of antigens. Specific binding of the target antibodies with their immobilized antigens on the surface results in a mosaic of binding events that can readily be visualized in one screening using fluorescence. It is thus possible to screen solns. for antibodies in a combinatorial fashion with great economy of reagents and at a high degree of miniaturization. Such mosaic-format immunoassays are compatible with the sensitivity and reliability required for immunodiagnostic methods. 27

REFERENCE COUNT:

THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 5 OF 20 CAPLUS COPYRIGHT 2002 ACS 1999:796050 CAPLUS

ACCESSION NUMBER:

132:20823 DOCUMENT NUMBER:

Detection of ligands with signal amplification TITLE: Woolverton, Christopher J.; Niehaus, Gary D.; INVENTOR(S):

Doane, Kathleen J.; Lavrentovich, Oleg; Schmidt,

Steven P.; Signs, Steven A. Kent State University, USA

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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APPLICATION NO.
                                                                                              DATE
                                  KIND DATE
       PATENT NO.
        ______
                                                                 WO 1999-US10413 19990512
                                            19991216
       WO 9964862
                                  A1
             W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM,
                    AZ, BY, KG, KZ, MD, RU, TJ, TM
             RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                                                   US 1998-95196
                                                                                              19980610
                                            20010109
                                   В1
       US 6171802
                                                                   AU 1999-39844
       AU 9939844
                                            19991230
                                                                                              19990512
                                    A1
                                                                   BR 1999-10982
                                                                                              19990512
       BR 9910982
                                    Α
                                            20010213
                                                                   EP 1999-922970
                                                                                              19990512
       EP 1086374
                                   Α1
                                            20010328
                   AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
                    PT, IE, FI
                                                               US 1998-95196
                                                                                          Α
                                                                                             19980610
PRIORITY APPLN. INFO.:
                                                              WO 1999-US10413 W 19990512
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A system for the detection of ligands comprising at least one AB receptor and an amplification mechanism coupled to the receptor wherein an amplified signal is produced as a result of receptor binding a ligand. Examples of suitable amplification mechanisms include antibody-embedded liq. cryst. materials; use of alpha-2-macroglobulin to encage an enzyme, whereby the enzyme is sepd. from its substrate by a receptor; and a receptor engineered to inhibit the active site of an enzyme only in the absence of a ligand. Also provided are methods for

the automatic detection of ligands.

THERE ARE 5 CITED REFERENCES AVAILABLE FOR REFERENCE COUNT: 5

THIS RECORD. ALL CITATIONS AVAILABLE IN

THE RE FORMAT

ANSWER 6 OF 20 CAPLUS COPYRIGHT 2002 ACS

1999:753249 CAPLUS ACCESSION NUMBER:

132:9589 DOCUMENT NUMBER:

Improved methods for detecting a . TITLE:

target nucleic acid fragment, particularly in

Borrelia burgdorferi and Babesia species

Shah, Jyotsna S.; Harris, Nick S. INVENTOR(S):

Igenex, Inc., USA PATENT ASSIGNEE(S): PCT Int. Appl., 50 pp. SOURCE:

CODEN: PIXXD2

Patent DOCUMENT TYPE: English LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

APPLICATION NO. DATE KIND DATE PATENT NO. ----_____ _____ ------_____ 19991125 WO 1999-US10939 19990518 WO 9960009 A1

W: AU, CA, JP

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,

NL, PT, SE

AU 9940011 A1 19991206 AU 1999-40011 19990518 US 1998-88541 P 19980521 PRIORITY APPLN. INFO.: P 19980521 US 1998-88696

WO 1999-US10939 W 19990518

The present invention provides a method for detecting a AB target nucleic acid fragment in a clin. specimen. A sample of the specimen is solubilized and treated to denature the nucleic acids therein. The sample is contacted with at least one probe complex comprising a sequence complementary to a portion of the target fragment, as well as a first member of a specific binding pair. Following hybridization of probe and target, the probe complex is contacted with a solid substrate linked to the second member of the specific binding pair to isolate hybridized target fragment in a probe-target-solid substrate ternary complex. The isolated ternary complex is sepd. from the sample, and the target fragment and probe complex are released into soln. The released target fragment is amplified by PCR or RT-PCR, and the presence of the target fragment in the clin. specimen is detected. The method may be used on a variety of specimen types, and is useful for detecting pathogens such as Borrelia burgdorferi and species of Babesia . Specific chaeotropic salt solns., wash buffers, and conditions for

amplification are disclosed. Preferred probe complexes and primers for the detection of Borrelia and Babesia

pathogens are disclosed.

REFERENCE COUNT:

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 7 OF 20 CAPLUS COPYRIGHT 2002 ACS 1999:41902 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 130:193412

Structure and mechanism of Ca2+-independent TITLE: phosphatidylinositol-specific phospholipases C Heinz, Dirk W.; Wehland, Jurgen; Griffith, O. AUTHOR(S): Hayes Institut fur Organische Chemie und Biochemie, CORPORATE SOURCE: Universitat Freiburg, Freiburg, D-79104, Germany ACS Symp. Ser. (1999), 718(Phosphoinositides), SOURCE: 80-90 CODEN: ACSMC8; ISSN: 0097-6156 American Chemical Society PUBLISHER: Journal; General Review DOCUMENT TYPE: LANGUAGE: English A review with 33 refs. The 3-dimensional structures of phosphatidylinositol-specific phospholipases C (PI-PLCs) from Bacillus cereus and the human pathogen, Listeria monocytogenes, have been detd. by x-ray crystallog ., both in free form and in complex with the substrate -like inhibitor, myo-inositol. Both enzymes share a very similar distorted (.beta..alpha.)8-barrel fold despite a moderate overall sequence identity of 24%. A high structural conservation has beenfound for the active site where myo-inositol is recognized in a stereospecific fashion. Two His residues that are also conserved between prokaryotic and eukaryotic PI-PLCs act as a general base and a general acid during catalysis, whereas an Arg residue provides the electrostatic stabilization of the transition state. Based on the present crystal structures and sequence alignments, it is suggested that all Ca2+-independent PI-PLCs known so far adopt the same fold and catalytic mechanism. THERE ARE 33 CITED REFERENCES AVAILABLE 33 REFERENCE COUNT: FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT ANSWER 8 OF 20 CAPLUS COPYRIGHT 2002 ACS 1998:757968 CAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 130:92086 The crystal structure of the L1 TITLE: metallo-.beta.-lactamase from Stenotrophomonas maltophilia at 1.7 .ANG. resolution Ullah, J. H.; Walsh, T. R.; Taylor, I. A.; AUTHOR(S): Emery, D. C.; Verma, C. S.; Gamblin, S. J.; Spencer, J. Division of Protein Structure, National CORPORATE SOURCE: Institute of Medical Research, London, NW7 1AA, UK J. Mol. Biol. (1998), 284(1), 125-136 CODEN: JMOBAK; ISSN: 0022-2836 SOURCE: Academic Press PUBLISHER: Journal DOCUMENT TYPE: English LANGUAGE: The structure of metallo-.beta.-lactamase L1 from the opportunistic pathogen, S. maltophilia, was detd. at 1.7 .ANG. resoln. by the multi-wavelength anomalous dispersion (MAD) approach exploiting both the intrinsic binuclear Zn center and incorporated selenomethionine residues. L1 is unique among all known .beta.-lactamases in that it exists as a tetramer. The protein was found to exhibit the .alpha..beta./.beta..alpha. fold found only in the metallo-.beta.-lactamases and displayed several unique features

Searcher: Shears 308-4994

not previously obsd. in these enzymes. These included a disulfide

bridge and 2 substantially elongated loops connected to the active site of the enzyme. Two closely spaced Zn2+ ions are bound at the active site with tetrahedral (Zn1) and trigonal bipyramidal (Zn2) coordination, resp.; these were bridged by a water mol. which it was proposed acts as the nucleophile in the hydrolytic reaction. Ligation of the 2nd Zn2+ ion involved both residues and geometry which had not been previously obsd. in the metallo-.beta.lactamases. Simulated binding of the substrates, ampicillin, ceftazidime, and imipenem, suggested that the substrate is able to bind to the enzyme in a variety of different conformations whose common features are direct interactions of the .beta.-lactam carbonyl O and N atoms with the Zn2+ ions and of the .beta.-lactam carboxylate with Ser-187. Here, the authors describe a catalytic mechanism whose principal features are a nucleophilic attack of the bridging water on the .beta.-lactam carbonyl C atom, electrostatic stabilization of a neg. charged tetrahedral transition state, and protonation of the .beta.-lactam N atom by a 2nd water mol. coordinated by Zn2. Further, the authors propose that direct metal-substrate interactions provide a substantial contribution to substrate binding and that this may explain the lack of specificity which is a feature of this class of enzyme. (c) 1998 Academic Press.

REFERENCE COUNT:

THERE ARE 47 CITED REFERENCES AVAILABLE 47 FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 9 OF 20 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER:

1996:614832 CAPLUS

DOCUMENT NUMBER:

125:241759

TITLE:

Crystal structures of Toxoplasma

gondii HGXPRTase reveal'the catalytic role of a

long flexible loop

AUTHOR(S):

Schumacher, Maria A.; Carter, Darrick; Roos, David S.; Ullman, Buddy; Brennan, Richard G. Dep. Biochem. Mol. Biol., Oregon Health Sciences

CORPORATE SOURCE:

Univ., Portland, OR, 97201-3098, USA

Nat. Struct. Biol. (1996), 3(10), 881-887

CODEN: NSBIEW; ISSN: 1072-8368

DOCUMENT TYPE:

Journal

LANGUAGE:

SOURCE:

English

Crystal structures of substrate-free and $\overline{\text{XMP-soaked}}$ hypoxanthine-guanine-xanthine phosphoribosyltransferase (HGXPRTase) of the opportunistic pathogen, T. gondii, were detd. to 2.4 and 2.9 .ANG. resoln., resp. HGXPRTase displayed the conserved phosphoribosyltransferase fold. In the structure of the enzyme bound to its product, a long flexible loop (residues 115-126) was located away from the active site. Comparison with the substrate-free structure revealed a striking relocation of the loop, which was poised to cover the catalytic pocket, thus providing a mechanism by which the HG(X)PRTases shield their oxocarbonium transition states from nucleophilic attack by the bulk solvent. The conserved Ser-117-Tyr-118 dipeptide within the loop was brought to the active site, completing the ensemble of catalytic residues.

ANSWER 10 OF 20 CAPLUS COPYRIGHT 2002 ACS 1.9

ACCESSION NUMBER:

1995:438161 CAPLUS

DOCUMENT NUMBER:

122:185337

308-4994 Searcher : Shears

TITLE:

y washing the sample area for a total of 6 min at 37.

A reading was taken of the reaction zone after the six minute time period and a second reading was taken 300 s later using a front surface fluorometer by directing 360 nm radiation through an opening in the assay module beneath the reaction zone and collecting the reflected 450 nm radiation. The increase in fluorescence, a function of the amt. of bound enzyme-labeled conjugate, was calcd. The result obtained was compared to the results obtained with a defined neg. calibrator and the pos. calibrators 15 described in Example Ill and was detd. thereby to be pos. or neg. on the basis of a detd. cutoff value.

L9 ANSWER 11 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1995:222764 CAPLUS

DOCUMENT NUMBER: 123:4385

TITLE: Crystal structure of scytalone

dehydratase -- a disease determinant of the rice pathogen, Magnaporthe

grisea

AUTHOR(S): Lundqvist, Tomas; Rice, Janet; Hodge, C

Nicholas; Basarab, Gregory S; Pierce, John;

Lindqvist, Ylva

CORPORATE SOURCE: Uppsala Biomedical Center, Swedish University of

Agricultural Sciences, Uppsala, S-751 24, Swed.

SOURCE: Structure (London) (1994), 2(10), 937-44

CODEN: STRUE6

DOCUMENT TYPE: Journal LANGUAGE: English

The 3-dimensional structure of scytalone dehydratase in complex with a competitive inhibitor was detd. at 2.9 .ANG. resoln. A novel fold, a cone-shaped .alpha.+.beta. barrel, was adopted by the monomer in this trimeric protein, burying the hydrophobic active site in its interior. The interactions of the inhibitor with the protein side chains were also identified. The similarity of the inhibitor to the substrate and the side chains involved in binding afforded some insights into possible catalytic mechanisms. These results provide a 1st look into the structure and catalytic

residues of a non-metal dehydratase, a large class of hitherto structurally uncharacterized enzymes. It is envisaged that a detailed structural description of scytalone dehydratase will assist in the design of new inhibitors for controlling rice blast disease.

L9 ANSWER 12 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1994:239633 CAPLUS DOCUMENT NUMBER: 120:239633

TITLE: Devices and methods for

detection of an analyte based upon light

interference

INVENTOR(S): Bogart, Gregory R.; Moddel, Garret R.; Maul,

Diana M.; Etter, Jeffrey B.; Crosby, Mark; Miller, John B.; Blessing, James; Kelley, Howard; Sandstrom, Torbjorn; Stiblert, Lars

PATENT ASSIGNEE(S): Biostar, Inc., USA

SOURCE: PCT Int. Appl., 208 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO.	KIND DATE			APPLICATION NO	٥.	DATE	
WO 9403774		19940217				19930610	
W: AT, AU, RW: AT, BE, SE	CA, JP CH, DE,	DK, ES,	FR, GI	B, GR, IE, IT,	LU	, MC, NL,	PT,
AU 9179004 AU 653940				AU 1991-79004		19910320	
EP 539383 EP 539383	A1 1	19930505		EP 1991-91005	6	19910320	
R: BE, CH,	DE, ES,	FR, GB,	IT, L	I, LU, NL, SE JP 1991-509344	4	19910320	
JP 05506936 JP 3193373 ES 2094224	B2 2	20010730		ES 1991-91005	6	19910320	
JP 3193373 ES 2094224 JP 2001235473 JP 07509565	A2 2	20010831		JP 2000-287242 JP 1993-505280	2	19910320 19930610	
EP 727038 R: ES FR.	GB. TT.	19960821 SE		EP 1993-91554.	ı	19930010	
EP 1126278 EP 1126278	A2 2	20010822		EP 2001-108523	1	19930610	
R: ES, FR, PRIORITY APPLN. INFO	CD TT	C F		1992-924343	A	19920731	
INIONITI MITEM. IMP	• •		, EP	1991-910056 1991-509344	1-7	19910320	
			WO	1991-US1781 1993-915341	Α	19910320	
			WO	1993-US5673	W	19930610	

Methods for analyzing an optical surface for an analyte of AB interest in a test sample and related instruments/devices are disclosed. The method entails the use of a thin-film optical immunoassay device whereby an analyte of interest is detected in a test sample through spectral changes in the light impinging on the surface prior to and after the binding of the analyte to a reactive substrate layer(s). The device includes a substrate which has a 1st color in response to light impinging thereon. The substrate also exhibits a 2nd color which is different from the 1st color. The 2nd color is exhibited in response to the same light when the analyte is present on the surface. Thus, SiO was vapor deposited on a polished monocryst. Si wafer to a thickness of 550 .ANG.; the film had a golden interference color. The film was activated with N-(2-aminoethyl)-3-aminopropyltrimethoxysilane, coated with aDNP-albumin conjugate to a thickness of 40.ANG., rinsed, and dried. The coated wafer was used in a competitive immunoassay for DNP using goat anti-DNP antibody and an ellipsometer to measure the change in mass at the surface from the change in light intensity.

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ANSWER 13 OF 20 CAPLUS COPYRIGHT 2002 ACS
                   1993:240872 CAPLUS
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ACCESSION NUMBER: 118:240872 DOCUMENT NUMBER:

Inhibition of mitochondrial respiration by TITLE:

furancarboxylic acid accumulated in uremic serum in its albumin-bound and non-dialyzable form

Niwa, T.; Aiuchi, T.; Nakaya, K.; Emoto, Y.; Miyazaki, T.; Maeda, K. AUTHOR(S):

CORPORATE SOURCE:

Dep. Intern. Med., Nagoya Univ., Nagoya, Japan

SOURCE:

Clin. Nephrol. (1993), 39(2), 92-6 CODEN: CLNHBI; ISSN: 0301-0430

DOCUMENT TYPE:

Journal English

LANGUAGE:

3-Carboxy-4-methyl-5-propyl-2-furanpropanoic acid (CMPF) accumulates markedly in uremic serum in its albumin-bound form. To det. if CMPF can be removed by newly developed dialyzers with high-flux membranes which are permeable to low-mol.-wt. proteins, such as .beta.2-microglobulin (.beta.2-MG), serum levels of CMPF were detd. before and after hemodialysis using these high-flux membrane dialyzers. In addn., to det. the pathogenic role of CMPF in uremic patients, its cellular toxicity due to its effect on mitochondrial respiration was studied. The redn. rates of CMPF by hemodialysis using the dialyzers ranged from -17% to -24%, demonstrating the nondialyzability of CMPF due to its strong

demonstrating the nondialyzability of CMPF due to its strong albumin-binding, while those of .beta.2-MG ranged from 11% to 43%. CMPF inhibited ADP-stimulated oxidn. of NADH-linked substrates in isolated mitochondria dose-dependently regardless of the presence of serum albumin. This inhibition was obsd. even at a concn. of 0.2 mM, which is comparable to the serum levels of CMPF in the hemodialysis patients. In conclusion CMPF

levels of CMPF in the hemodialysis patients. In conclusion CMPF which cannot be removed even by high-flux membrane dialyzers, is a strong inhibitor of mitochondrial respiration, and novel purifn. methods to remove CMPF from the blood of uremic patients

should be developed.

9 ANSWER 14 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1992:169623 CAPLUS

116:169623

DOCUMENT NUMBER: TITLE:

Substrate-adhered polymyxin

immunoassay device for

lipopolysaccharide testing, and method

for its preparation

INVENTOR(S):

Blais, Berton W.; Yamazaki, Hiroshi

PATENT ASSIGNEE(S):

Can.

SOURCE:

Can. Pat. Appl., 54 pp.

CODEN: CPXXEB

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

IND	DATE	APPLICATION NO.	DATE
AA	19911118	CA 1990-2017093	19900518
A2	19920928	JP 1991-113467	19910517
A	19960423	US 1993-87013	19930707
	CA	1990-2017093	19900518
	CA	1991-2037726	19910307
	CA	. 1991-2037727	19910307
	US	1991-697683	19910509
	AA A2	AA 19911118 A2 19920928 A 19960423 CA CA	AA 19911118 CA 1990-2017093 A2 19920928 JP 1991-113467

AB An improved ap. is disclosed for the detection of Gram-neg. bacterial lipopolysaccharide (LPS). The improved app. and process uses a min. of expensive antibody. Polymixin is adhered to a substrate of e.g. plastic or (non)woven cloth. The Polymyxin can then bind to LPS which may in turn be identified by a std. immunoassay. Nonwoven polyester cloth is esp.

suited for the device. The app. is esp. useful for the detection of bacterial pathogens in food. A polymyxin B-coated nonwoven polyester cloth was prepd. and used in the detection of Salmonella.

ANSWER 15 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1991:225216 CAPLUS

DOCUMENT NUMBER: 114:225216

Parallel solid-phase method to TITLE:

determine multiple immunologically detectable

substances

Bayer, Hubert; Kirch, Peter; Kopetzki, Erhard; INVENTOR(S):

Klein, Christian

Boehringer Mannheim G.m.b.H., Fed. Rep. Ger. PATENT ASSIGNEE(S):

Eur. Pat. Appl., 12 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE:

Patent German LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 379216	A1	19900725	EP 1990-101095	19900119
EP 379216	B1	19940608	EL 1930 101030	
	E, CH, DE	, DK, ES,		
DE 3901638	A1	19891207	DE 1989-3901638	19890120
DE 3901638	C2	19990325	DĖ 1989-3924239	19890721
DE 3924239 AT 107030	A1 E	19910124 19940615	AT 1990-101095	19900119
	FO.:	19940013	DE 1989-3901638 A	19890120
			DE 1989-3924239 A	19890721
			DB 1300 001. 120 112	19880525
			EP 1990-101095 A	19900119

A parallel solid-phase immunoassay or specific binding assay for detg. multiple analytes in a sample (e.g. multiple antibodies against different epitopes on a virus) uses (1) a specific binding partner (e.g. avidin) bound to a solid phase; (2) a series of receptors R1, each comprising a conjugate of the complementary specific binding partner (e.g. biotin) and a ligand for 1 of the analytes; and (3) a series of receptors R2, each comprising a conjugate of a ligand for 1 of the analytes and a detectable moiety (label). The ligand and label on receptors R2 may be the same for all analytes (allowing e.g. the detection of a virus in all its variants) or may be different for each analyte (allowing the detection of each analyte individually). streptavidin was coupled to bovine serum albumin via maleimidohexanoyl-N-hydroxysuccinimide and adsorbed on the surface of a polystyrene tube. In a test for antibodies to human immunodeficiency virus (HIV), the tube was incubated with .gtoreq.1 biotinylated HIV antigen and a serum or plasma sample, washed, incubated with a a sheep anti-human Ig antibody conjugated to peroxidase, washed, and incubated with a peroxidase substrate (ABTS) for photometric detection.

ANSWER 16 OF 20 CAPLUS COPYRIGHT 2002 ACS

1991:120157 CAPLUS ACCESSION NUMBER: 114:120157 DOCUMENT NUMBER:

> 308-4994 Searcher : Shears

TITLE: Avidin-biotin-assisted immunoassay using

immunoaffinity chromatography for antibody or

antigen detection

INVENTOR(S): Thieme, Thomas; Ferro, Adolph; Fellman, Jack H.;

Gavojdea, Stefan Epitope, Inc., USA

PATENT ASSIGNEE(S): Epitope, Inc., USA SOURCE: PCT Int. Appl., 26 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA'	rent :	NO.		KII	ND	DATE			Al	PPLI	CATIO	ои ис	٥.	DATE		
MO.	9008											s378		1990		
	W:	ΑU,	BB,	BG,	BR,	CA,	FΙ,	ΗU,	JΡ,	KP,	KR,	LK,	MC,	MG,	MW,	NO,
			SD,													
	RW:	ΑT,	ΒE,	BF,	ВJ,	CF,	CG,	CH,	CM,	DE,	DK,	ES,	FR,	GA,	GB,	IT,
		LU,	ML,	MR,	NL,	SE,	SN,	TD,								
CA	2045	673					0731			_		0456		1990		
AU	9050	986					0824		ΑŪ	J 19	90-50	0986		1990	0125	
AU	6406	35					0902						_			
EP	4557	35					1113		El	P 19	90-9	0346	6	1990)125	
EP	4557					1994										
	R:	AT,	BE,	DE,			GB,						_			
	0450				_		0604					0351		1990		
NO	9102	951		Α		1991	0913				91-2			1991	-	
PRIORIT	Y APP	LN.	INFO	.:				1	US 19	989-	3028	77		1989		
								1	WO 19	990-1	US37	8		1990	0125	

A method is provided for antibody detection in a body AB fluid for screening and diagnostic purposes. The method uses an antibody-binding protein bonded to a porous matrix within a transparent column. A test fluid contg. the antibody is contacted with the protein-bonded matrix to immobilize the antibody. A biotinylated antigen of interest is contacted with the immobilized antibody, and enzyme-linked avidin is contacted with the resulting immobilized complex. This final immobilized complex is contacted with enzyme substrate to produce a colored product which correlates to the presence of the antibody to be detected. Color comparison controls may be run in the same transparent column using addnl. aliquots of agarose sepd. by polyethylene disks. Thus, the above method was used with protein A-agarose matrix, avidin-peroxidase conjugate, and biotinylated human immunodeficiency virus (HIV) lysate to detect antibodies against HIV-1 in serum and saliva of AIDS patients. Using antibody-pos. samples, a near max. dark brown color was obtained on the agarose beads following the enzymic color development reaction. Control serum and saliva gave essentially colorless agarose beads. The **methods** of the invention are also used for antigen detection.

L9 ANSWER 17 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1990:173609 CAPLUS

DOCUMENT NUMBER: 112:173609

TITLE: Colorimetric detection of PCR products using the

DNA-binding protein TyrR

AUTHOR(S): Triglia, Tony; Argyropoulos, Victor P.;

Davidson, Barrie E.; Kemp, David J.

Walter and Eliza Hall Inst. Med. Res., CORPORATE SOURCE:

Melbourne, 3050, Australia

Nucleic Acids Res. (1990), 18(4), 1080 SOURCE:

CODEN: NARHAD; ISSN: 0305-1048

DOCUMENT TYPE: Journal LANGUAGE: English

Recent methods eliminating the need for electrophoresis in the detection of PCR products include the use of the DNA-

binding protein, GCN4 or magnetic beads. The DNAbinding protein, TyrR, an Escherichia coli protein

responsible for the regulation of expression of eight operons

involved in arom. biosynthesis and transport, can be used much like

GCN4. Basically, the method involves binding

GCN4 or TyrR to the wells of microtiter plates, then adding PCR-amplified DNA which has either a GCN4 or a TyrR binding

site attached to one oligonucleotide and a biotin group attached to

the other oligonucleotide. Simultaneously, avidin-labeled peroxidase binds to the biotin. After washing, the

peroxidase can be detected easily with a chromogenic substrate. Only GCN4 (and not TyrR) coated wells

bind the PCR-amplified material with the GCN4

binding site, and only the TyrR-coated wells bind the PCR-amplified material with the TyrR binding site.

The use of two different DNA-binding proteins should allow

the simultaneous detection of two different pathogens or two genotypes of the same pathogen.

ANSWER 18 OF 20 CAPLUS COPYRIGHT 2002 ACS

1989:532410 CAPLUS ACCESSION NUMBER: DOCUMENT NUMBER:

111:132410

TITLE:

Antigenic determinants recognized by antibodies

obtained using a pathogenic agent or a

derivative thereof that presents a restricted set of antigens, and production of antimalaria

vaccines

INVENTOR(S): Lyon, Jeffrey A.; Chulay, Jeffrey D.; Thomas,

Alan W.; Howard, Russel J.; Weber, James L.

United States Dept. of the Army, USA PATENT ASSIGNEE(S):

SOURCE:

PCT Int. Appl., 50 pp.

CODEN: PIXXD2

Patent DOCUMENT TYPE:

LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE PATENT NO. KIND DATE WO 1988-US788 19880314 19880922 WO 8806892 A1

W: JP

RW: AT, BE, BJ, CF, CG, CH, CM, DE, FR, GA, GB, IT, LU, ML, MR,

NL, SE, SN, TD, TG

US 1987-25741 19870313 US 4906564 19900306 Α A1 19890315 EP 1988-903541 19880314 EP 306524

R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE

19870313 PRIORITY APPLN. INFO.: US 1987-25741

A method is presented for identifying antigenic determinants of a pathogenic agent, comprising (A)

providing a sample of intact pathogen, (B) contacting the sample

Shears 308-4994 Searcher :

with a heterogeneous mixt. of antibodies from functional immune serum, such that an antibody from the mixt. binds to an antiquenic determinant on the pathogen, (C) isolating the antibody from the sample, and (D) using the antibody to probe a heterogeneous mixt. of antigens produced by the pathogen. Antigens so identified are used in vaccines against the pathogen. Plasmodium falciparum schizont-infected erythrocytes were cultured for 5 h at 37.degree. in a functional immune serum from Aotus monkeys; during this time, the merozoites released from the infected erythrocytes developed a thick coat of antigen-antibody complexes and became agglutinated. These merozoites were extd. with Triton X-100 at pH 8 to obtain antibodies. DNA from P. falciparum merozoites was ligated into the lacZ gene of .lambda.gtll vector DNA for prodn. of a gene library in Escherichia coli. Phage plaques were overlayed with a IPTG-treated nitrocellulose filter to induce expression of the lacZ gene. Then the antibody prepn. was applied to the filter, followed by washing, application of rabbit anti-Aotus IgG conjugated to alk. phosphatase, and application of chromogenic substrate. Of 80,000 plaques adsorbed on the filter, 20 showed color formation indicative of antigen-antibody complexing; 5 of these were cloned and found to code for merozoite surface glycoprotein gp195.

L9 ANSWER 19 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1987:100758 CAPLUS

DOCUMENT NUMBER: 106:100758

TITLE: Method and reagent kit for

pathogenic streptococcal antibody

determination

INVENTOR(S): Hayano, Seiki; Yokogawa, Kanae; Kurooka, Shigeru

PATENT ASSIGNEE(S): Dainippon Pharmaceutical Co., Ltd., Japan SOURCE: U.S., 7 pp. Cont. of U.S. Ser. No. 249,231,

abandoned.
CODEN: USXXAM

DOCUMENT TYPE: Patent

type A-I esterase antibody.

LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

APPLICATION NO. DATE PATENT NO. KIND DATE -----_____ _____ US 1983-552014 US 4592995 19860603 19831116 Α US 1981-249231 19810330 PRIORITY APPLN. INFO.: Pathogenic streptococcal esterase antibody in human blood serum is quant. detd. by (a) adding a pathogenic streptococcal esterase to the blood sample; (b) adding an immobilized protein-A which nonspecifically binds to the antibody to form an esterase-antibody-insol. protein-A complex; (c) sepg. the complex from the mixt. by centrifugation; and (d) measuring the activity of the esterase in the complex. The kit and method are useful for diagnosis of various diseases caused by pathogenic streptococcal infections. Type A-I esterase, prepd. from Streptococcus pyogenes, was added to dild. patient serum and incubated. Cell walls of Staphylococcus aureus Cowan I, contg. protein-A, were added next and incubated. The reaction mixt. was washed and centrifuged. The esterase activity was measured at 412 nm after addn. of S-acetylthiophenol as substrate to det.

L9 ANSWER 20 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1985:42494 CAPLUS

DOCUMENT NUMBER: 102:42494

TITLE: Methods and structures employing non-radioactive chemically-labeled

Patent

polynucleotide probes

INVENTOR(S): Stavrianopoulos, Jannis G.; Kirtikar, Dollie;

Johnston, Kenneth H.; Thalenfeld, Barbara E.

PATENT ASSIGNEE(S): Enzo Bio Chem, Inc., USA SOURCE: Eur. Pat. Appl., 76 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	PATENT NO.		KIND DATE		APPLICATION NO.	DATE
	117440 117440		A1	19840905 19930407	EP 1984-100836	19840126
					IT, LI, LU, NL, SE	
IL	70765		A1		IL 1984-70765	19840123
CA	1309672		A1	19921103	CA 1984-445896	19840123
DK	8400313				DK 1984-313	19840124
NO	8400289		Α	19840730	NO 1984-289	19840125
AU	8423798		A1	19840802	AU 1984-23798	19840126
AU	577776		В2	19881006		
ES	529179		A1	19850616	ES 1984-529179	19840126
EP	525821		A2	19930203	EP 1992-114727	19840126
EP	525821		ΑЗ	19940302		
	R: AT,	BE,	CH, DE,	FR, GB,	IT, LI, LU, NL, SE	
	87929		E	19930415	AT 1984-100836	19840126
JP	59141599		A2	19840814	JP 1984-14165	19840127
JP	2825090		В2	19981118		
ES	540485		A1	19860216	ES 1985-540485	19850216
US	4994373		A	19910219	US 1989-385986	19890720
	Y APPLN.				US 1983-461469	19830127
•					EP 1984-100836	19840126
					US 1985-732374	19850509
			,		and the state of DMA completes and	

AB The title probes, such as single-stranded DNA probes contg.
.gtoreq.25 bases, contain esp. enzyme labels and are used for the detection and identification of complementary single-stranded DNA (fixed on an inert support) by hybridization, followed by spectrophotometric detn. of the enzyme in the double-stranded hybrids formed. These probes can replace hazardous, expensive, short-lived radiolabeled probes, and ELISA also can be used to det. the formed hybrid. The support may be glass, polystyrene, nitrocellulose, dextran, etc. Recommended enzyme labels and some substrates are tabulated. The method is useful for detecting the presence of pathogens (e.g., Streptococcus, Staphylococcus, Pneumococcus, etc.) in clin. samples by detection of their genetic material.

LUS, JAPIO' ENTERED AT 15:34:17 ON 06 MAR 2002)

15 S L3 47 S L8

L10 L11

L12

58 S L10 OR L11

47 DUD DEM 112 (11 DUPLICATES REMOVED)

L13 ANSWER 1 OF 4/ACCESSION NUMBER:

L13 ANSWER 1 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

2002-083189 [11] WPIDS

DOC. NO. CPI:

C2002-025293

TITLE:

Analyzing variant sites of target nucleic acids,

useful for identifying and detecting point

mutations, specifically those mutations correlated

with diseases e.g. cancer, by limited primer

extension.

DERWENT CLASS:

B04 D16

INVENTOR(S):
PATENT ASSIGNEE(S):

GLAZER, A N; XU, H (DNAS-N) DNA SCI INC

COUNTRY COUNT:

94

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001092583 A1 20011206 (200211)* EN 54

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE

DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG

KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ

PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 200109258	3 A1	WO 2001-US18023	20010531

PRIORITY APPLN. INFO: US 2000-586125 20000602

AN 2002-083189 [11] WPIDS

AB WO 200192583 A UPAB: 20020215

NOVELTY - Analyzing (M1) a variant site of a target nucleic acid (TNA) by conducting template dependent extension reaction in presence of a mixture of labeled extendible nucleotide (nt) (LEN) and labeled non-extendible nts (LNEN), and detecting incorporation of labeled nt indicating identity of nt at variant site, since incorporated nt is complementary to nt at site of variation.

DETAILED DESCRIPTION - Analyzing (M1) a variant site of a target nucleic acid (TNA) by conducting template dependent extension reaction in presence of a mixture of labeled extendible nucleotide (nt) (LEN) and labeled non-extendible nts (LNEN), and detecting incorporation of labeled nt indicating identity of nt at variant site, since incorporated nt is complementary to nt at site of variation. (M1) comprises:

(a) conducting a template-dependent extension reaction comprising extending a primer (I) in the presence of TNA and a mixture of nucleotides comprising a LEN and LNEN being complementary to a different allelic form of TNA and optionally differentially labeled, where (I) hybridizes to a segment of TNA such that the 3'-end of (I) hybridizes adjacent to the variant site of TNA, where if the LEN is complementary to the nucleotide occupying the variant

- site, (I) is extended by incorporation of the LEN, and can be extended further if one or more nucleotides downstream of the variant site are complementary to one of the nucleotides in the mixture, and if the LNEN is complementary to the nucleotide occupying variant site, (I) is extended by incorporation of the LNEN; and
- (b) detecting incorporation of labeled nucleotide into the extended (I), the identity of the labeled nucleotide incorporated into (I) indicating the identity of the nucleotide at the variant site, where the identity of the incorporated nucleotide is determined from the label borne by the incorporated nucleotide and/or the size of the extended (I).

INDEPENDENT CLAIMS are also included for the following:

- (1) analyzing (M2) variant sites in one or more TNA comprising:
- (a) conducting several template-dependent extension reactions in the presence of several different primers, where the primers hybridize adjacent to different variant sites of TNAs and are differentially labeled, where extension reaction comprises contacting a sample containing the TNAs with one of the different labeled primers, and exposing the primer to a mixture of nucleotides comprising LEN and LNEN, where the extension reactions generate several different extension products, which are from different variant sites being distinguishable on the basis of the different labels borne by the extended primers; and
- (b) detecting incorporation of labeled nucleotides into the extension products as an indication of the nucleotides occupying the site of variation in TNAs, where the identity of the incorporated nucleotide is determined from the label borne by the incorporated nucleotide and/or the size of the extended primer; and
- (2) a kit (II) utilized in (M1) comprising LEN , LNEN, nucleotides complementary to different allelic forms of TNA, and a primer that hybridizes to a segment of TNA such that 3' end of the primer adjacent to the variant site of TNA.

USE - M1 is useful for analyzing a variant site of a target nucleic acid, and M2 is useful for analyzing variant sites in one or more TNA (claimed).

The method and (II) is useful for identification and detection of point mutations (e.g. somatic point mutation), specifically those mutations correlated with diseases such as diseases associated with SNPs which include sickle cell anemia, cystic fibrosis; autoimmune diseases; formation of oncogenes and cancer. For e.g. identifying whether a nucleic acid from a particular subject includes a wild-type allele or a mutant allele at a particular single nucleotide polymorphic (SNP) site. Further, the methods can be utilized to establish the genotype of the individual being tested (i.e., distinguish whether the individual is a reference-type homozygote, a heterozygote or a variant-type homozygote). The genotyping utility of the methods makes them useful within the context of medical diagnosis and prognosis. Since many SNPs are associated with various diseases and clinicians can utilize the results of the genotype study to assess the presence of disease, whether an individual is a carrier of disease, the likelihood that an individual will get a particular disease and the likely efficacy of various treatment alternatives.

The methods also have a variety of non-medical uses, such as detecting pathogenic microorganisms, paternity testing and forensic analysis in which polymorphisms in specific games can be determined in, for e.g. blood or semen

obtained from a crime scene to indicate whether a particular suspect was involved in the crime. In like manner, polymorphism analysis may be utilized in disputes to aid in determining whether a particular individual is the parent of a certain child. The **methods** can also be used to identify SNPs in non-humans, including, for e.g. other animals, plants, bacteria and viruses.

The methods are also useful for identifying point mutations in pathogens that could potentially result in altered pathogenicity or resistance to certain therapeutics; and to identify cells and strains having a desired genetic constitution for use in various biotechnology applications. The method is utilized as a diagnostic tool and a prognostic tool of a disease which is useful in formulating optimal treatment for the patient.

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L13 ANSWER 2 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2002-114152 [15] WPIDS

DOC. NO. NON-CPI: N2002-085150 DOC. NO. CPI: C2002-034959

TITLE: Analysis of polynucleotides in a sample using

generic capture sequences comprises amplifying target polynucleotide, and utilizing the product to indirectly assay the sample for the polynucleotide.

DERWENT CLASS: B04 D16 S03

INVENTOR(S): LAI, J H; PHILLIPS, V E; WATSON, A R

PATENT ASSIGNEE(S): (QUAN-N) QUANTUM DOT CORP

COUNTRY COUNT: 95

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001083823 A1 20011108 (200215)* EN 85

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ

DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE
WO 2001083823 A1 WO 2001-US13979 20010430

PRIORITY APPLN. INFO: US 2000-200635P 20000428

AN 2002-114152 [15] WPIDS

AB WO 200183823 A UPAB: 20020306

NOVELTY - Assaying, (M1) for an amplification product (AMP) from a first target polynucleotide, (TP), comprising providing a sample that is suspected of containing AMP, where the AMP is a polynucleotide comprising a first label and a capture sequence not present in the TP at the same position, is new.

DETAILED DESCRIPTION - A target polynucleotide, (TP), is amplified, where first primer has a tag sequence, the complement of which is formed on the opposite strand during amplification and is

referred to as capture sequence (CS), and the opposite strand is referred to as an amplification product (AMP) has a label; probe conjugated to substrate specific to CS, is contacted with AMP to form AMP detection complex. Assaying (M1) for an AMP from a first TP, comprises providing a sample that is suspected of containing AMP, where the AMP is a polynucleotide comprising a first label and a capture sequence not present in the TP at the same position, where the AMP is formed by primer extension from a template, where the template comprises a complement to the TP and a target noncomplementary region, where the capture sequence is a complement to the target noncomplementary region, providing a substrate that is conjugated to a first capture probe, contacting the sample with the capture probe under a first set of hybridization conditions, where the capture probe can bind to the capture sequence under the first set of hybridization conditions and determining if the first label is associated with the substrate.

INDEPENDENT CLAIMS are also included for the following:

- (1) forming (M2) an AMP detection complex for assaying a sample for a first TP;
- (2) an amplification product detection complex comprising a capture probe polynucleotide hybridized to capture sequence of labeled amplification product from a TP, where the capture probe polynucleotide is conjugated to a substrate, where the capture sequence is not present in a region of the TP which is amplified and is introduced into the amplification product by copying a template polynucleotide, the template polynucleotide comprising a target noncomplementary region and a region complementary to the TP, where the target noncomplementary region was introduced into the template polynucleotide by extension of a primer hybridized to the target polynucleotide, the primer comprising the target noncomplementary region and where the capture sequence is complementary to the target noncomplementary region; and
- substrate attached to a capture probe, a first primer comprising a 3' end a first target complementary region located at the 3' end of the first primer, and a first target noncomplementary region that is not complementary to the first TP at a position 3' of a sequence to which the first target complementary region can hybridize, a second primer, label, housing for retaining the substrate, first primer, second primer, and a label and instruction provided with the housing that describe how to use the components of the kit to assay a sample by forming an AMP from the TP using the first and second primers that comprises the label and a capture sequence that is complementary to the target noncomplementary in the first primer, where the capture sequence can bind to the capture probe.
- USE (M2) is useful for forming an AMP detection complex for assaying a sample for first TP. The method further comprises determining if the first label is associated with the first substrate, where AMP is produced at a detectably higher level from at least one allele of a locus having at least two alleles and the first substrate preferably a first microsphere comprising a first spectral code is identified by decoding the spectral code which is preferably performed prior to, simultaneously or subsequent to determining if the first label from the second primer is associated with a substrate, first TP, preferably single-stranded or double-stranded DNA or RNA and a

polymerase, preferably DNA polymerase having reverse transcriptase activity is used to form the first primer extension product (claimed). (M1) is useful for particular polynucleotide sequences, whether based on SNPs, conserved sequences, or other features or useful in a wide variety of different applications. The method is useful for pharmacogenetic testing, such methods ca be used in a forensic setting to identify the species or individual which was the source of a forensic specimen. Polynucleotide analysis methods can also be used in an anthropological setting. Paternity testing is another area, as is testing for compatibility, between prospective tissue or blood donors and patients in need, and in screening for heredity disorders. (M1) is also useful for studying gene expression in response to a stimulus. Other applications include human population genetics, analyses of human evolutionary history, and characterization of human haplotype diversity. The method is useful to detect immunoglobulin class switching and hypervariable mutation of immunoglobulins, to detect polynucleotide sequences from contaminants or pathogens including bacteria, yeast, viruses, for HIV subtyping to determine the particular strains or relative amounts of particular strains infecting an individual, and can be repeatedly to monitor changes in the individuals predominant HIV strains, such as the development of drug resistance or T cell tropism; and to detect single nucleotide polymorphisms, which may be associated with particular alleles or subsets of allele. Over 1.4 million different single nucleotide polymorphisms (SNPs) in the human population have been identified. The method is also useful for mini-sequencing and for detection of mutations. Any type of mutation can be detected, including without limitation SNPs, insertions, deletions, transitions, transversions, inversions, frame shifts, triplet repeat expansions, and chromosome rearrangements. The method is useful to detect nucleotide sequences associated with increased risk of diseases or disorders, including cystic fibrosis, Tay-Sachs, sickle-cell anemia, etc. The method is useful for any assay in which a sample can be interrogated regarding an amplification product from a target polynucleotide. Typical assays involve determine the presence of the amplification product in the sample or its relative amount, or the assays may quantitative or semi-quantitative. Results from such assays can be used to determine the presence or amount of the target polynucleotide present in the sample. The above methods are particularly useful in multiplex settings where several TP are to be assayed. Dwg.0/15

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L13 ANSWER 3 OF 47 WPIDS COPYRIGHT 2002
                                           DERWENT INFORMATION LTD
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2002-114149 [15] WPIDS ACCESSION NUMBER:

N2002-085149 DOC. NO. NON-CPI:

C2002-034956 DOC. NO. CPI:

Microfabricated electrochemical biosensor, useful TITLE: for detecting ions, proteins and nucleic acid,

produced by integrated circuit technology.

B04 D16 J04 L03 S03 S05 U12 DERWENT CLASS:

GAU, J INVENTOR(S):

(GAUJ-I) GAU J PATENT ASSIGNEE(S):

95 COUNTRY COUNT:

PATENT INFORMATION:

308-4994 Shears Searcher :

PATENT NO KIND DATE WEEK LA PG

WO 2001083674 A1 20011108 (200215)* EN 84

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ

DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE

KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO

NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ

VN YU ZA ZW

APPLICATION DETAILS:

PRIORITY APPLN. INFO: US 2000-201603P 20000503

AN 2002-114149 [15] WPIDS

AB WO 200183674 A UPAB: 20020306

NOVELTY - Detecting or quantifying an analyte (I) by applying a sample to the electrodes of a microfabricated electrochemical biosensor and measuring an electrical signal, is new. The biosensor comprises a substrate having fabricated on it, by integrated circuit (IC) technology, at least two electrodes that consist of a single layer of conductive material.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for

the following:

- (1) microfabricated electrochemical biosensor comprising a silicon substrate and three redox-sensing electrodes, each a single layer of gold formed by IC technology;
- (2) detecting (I) by applying a sample to a biosensor having two surface areas with different properties, one for immobilizing (I) and the other for:
- (a) confining the sample by surface tension forces between the two areas; and
 - (b) for detecting (I); and
- (3) **device** for detecting a redox event of at least one (I).

USE - For detecting/determining ions (metals) or macromolecules (DNA, RNA or proteins), e.g. in rapid detection of pathogenic bacteria.

ADVANTAGE - The sensor may incorporate hybridization and enzymatic amplification for increased sensitivity and miniaturization. Sample and reagent can be confined by surface tension forces, making it possible to incorporate the sensor in portable/hand-held instruments and protecting them against shaking or inversion. The entire sensor system can be prepared on a single chip, eliminating the need for external components. The sensors are easy to produce, inexpensive and reusable, with the same robustness and reversible electrochemical performance as conventional sensors. By confining sample and detection reagent to specific regions of the substrate, problems of non-specific binding are overcome (improving sensitivity), only very small samples are required (ensuring transport by diffusion, without the need for any mixing), coverage of the electrodes is easily controlled and loss of target analyte is minimized.

Dwg.0/35

L13 ANSWER 4 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2002-017643 [02] WPIDS

DOC. NO. NON-CPI: N2002-014058 DOC. NO. CPI: C2002-005119

TITLE: A new spatially defined array comprises protein expression systems bound at discrete positions to a

substrate and is useful to screen for

potentially useful compounds such as receptor

ligands.

DERWENT CLASS: BO4 D16 S03 T01

INVENTOR(S): PATRON, A; SAWAFTA, R; ZHOU, B

PATENT ASSIGNEE(S): (PATR-I) PATRON A; (SAWA-I) SAWAFTA R; (ZHOU-I)

ZHOU B; (TRAN-N) TRANS TECH PHARMA

COUNTRY COUNT: 94

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001079849 A2 20011025 (200202)* EN 21

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

US 2001041349 A1 20011115 (200202)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 20010798 US 20010413	49 A2 49 A1 Provisional	WO 2001-US12474 US 2000-197692P US 2001-836746	20010417 20000417 20010417

PRIORITY APPLN. INFO: US 2000-197692P 20000417; US 2001-836746 20010417

AN 2002-017643 [02] WPIDS

AB WO 200179849 A UPAB: 20020109

NOVELTY - A spatially defined array (I) of protein expression systems, comprising a substrate, a binding

surface which covers some or all of the **substrate** surface and a number of protein expression systems located at discrete positions on portions of the **substrate** covered by the **binding** surface, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- a micromachined device comprising (I);
- (2) a biosensor comprising the claimed array;
- (3) screening (M1) a number of proteins for their ability to interact with a component of a sample, comprising generating the claimed array and detecting interaction of the component with proteins expressed at specific positions comprising the protein expression systems;

(4) detecting (M2) chemical or biological components immobilized on a solid phase by multidimensional spectroscopy using ion mobility and time of flight mass spectroscopy, comprising:

(a) recovering at least part of a chemical or biological mixture immobilized on a solid substrate as an electrospray;

(b) directing the electrospray to an ion mobility chamber which separates the constituents of the spray by size, ionic charge and

(c) analyzing the separated constituents which emerge from the chamber by time of flight spectroscopy; and

(5) computer-readable media comprising software code for performing the above method.

USE - The invention is used in biochemical research to screen new compound such as potential receptor ligands and small molecules.

ADVANTAGE - Unlike prior art arrays the protein does not need

to be purified for array generation.

DESCRIPTION OF DRAWING(S) - Figure shows the array of the invention used in sequestering proteins. Panel A shows host cells expressing a soluble protein. B shows host cells expressing a protein with an affinity tag. C shows host cells expressing a membrane bound protein.

soluble protein 40

cells 42

wells 46,48

affinity tags 50

binding partner 42 expression array 46,48

Dwg.3/4

L13 ANSWER 5 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

2002-097393 [13] WPIDS ACCESSION NUMBER:

N2002-071996 DOC. NO. NON-CPI:

DOC. NO. CPI: C2002-030253

TITLE: New integrated microscale biosensor that is microfluidic system with integrated elements, inlet-outlet ports and interface schemes, useful

for detecting pathogens, e.g. foodborne Listeria monocytogenes.

DERWENT CLASS: A89 B04 D16 S05

BASHIR, R; BHUNIA, A K; GOMEZ, R; LADISCH, M R; INVENTOR(S):

ROBINSON, J P; SARIKAYA, A

(PURD) PURDUE RES FOUND PATENT ASSIGNEE(S):

COUNTRY COUNT: 95

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA

WO 2001079529 A1 20011025 (200213)* EN 81

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO

NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW

US 2001053535 A1 20011220 (200213)

APPLICATION DETAILS:

PATENT NO KI	ND	API	PLICATION	DATE
WO 2001079529 US 2001053535		US	2001-US9745 2000-197560P 2001-817541	20010326 20000417 20010326

PRIORITY APPLN. INFO: US 2000-197560P 20000417; US 2001-817541

20010326

AN 2002-097393 [13] WPIDS

AB WO 200179529 A UPAB: 20020226

NOVELTY - A biosensor or integrated microscale biosensor, comprising a substrate microfabricated to include as integrated components detection chambers and channels, is new.

DETAILED DESCRIPTION - A biosensor or integrated microscale biosensor, comprising a **substrate** microfabricated to include as integrated components:

- (a) a detection chamber;
- (b) a first channel segment extending to an inlet of the detection chamber;
- (c) a second channel segment extending from an outlet of the chamber; and
- (d) a retention structure for holding, in the chamber, carrier elements entraining a target microbiological species while permitting passage through the detection chamber of contaminant materials in a fluid stream.

The integrated microscale biosensor may comprise a substrate microfabricated to include as integrated components:

- (a) a detection chamber;
- (b) a channel extending to an inlet of the detection chamber; and
- (c) an inlet groove or trench substantially coplanar with the channel and the detection chamber, further comprising an elongate fluid delivery member having a downstream end disposed in the inlet groove or trench, the fluid delivery member being connected at the downstream end to inlet groove or trench so that at least the downstream end of the fluid delivery member is coplanar with the channel and the detection chamber.

INDEPENDENT CLAIMS are also included for the following:

- (1) detecting a microbiological substance comprising:
- (a) providing the microfabricated biosensor chip including integrated detection elements;
 - (b) delivering a fluid sample to the biosensor chip;
- (c) after the delivering of the fluid sample to the biosensor chip, separating at least some contaminants from the fluid sample to at least partially isolate and retain instances of a predetermined type of microbiological material on the biosensor chip, the separating of the contaminants taking place on the biosensor chip; and
- (d) after the separating of contaminants from the fluid sample, operating the detection elements to determine whether the separated fluid sample contains microbiological material of the predetermined type;
 - (2) manufacturing a biosensor comprising:
 - (a) providing a substrate;

- (b) processing the substrate to generate a detection chamber and a channel extending to the detection chamber;
- (c) further processing the substrate to provide at least one pair of electrodes in the detection chamber; and
- (d) exposing the processed **substrate** to bovine serum albumin (BSA) and avidin to adsorb the avidin to the electrodes in the presence of the BSA;
 - (3) manufacturing a biosensor comprising:
 - (a) providing a substrate;
- (b) processing the substrate to create a shallow detection chamber and a channel extending to the detection chamber;
- (c) after the creation of the detection chamber and the channel, further processing the substrate to deposit at least one pair of electrodes in the detection chamber;
- (d) after the deposition of the electrodes, further processing the substrate to create at lest one deep groove at a periphery of the substrate, for receiving an elongate fluid delivery element, the channel communicating with the deep groove;
- (e) inserting a downstream end of the fluid delivery element into the deep groove; and
- (f) attaching the downstream end of the fluid delivery element to the deep groove;
 - (4) detecting a microorganism comprising:
- (a) preparing a fluid sample containing at least one microorganism of a preselected type, the fluid sample having a buffer of a low conductivity liquid, the fluid sample also containing a nonionic nutrient;
- (b) disposing the fluid sample in a detection chamber having a volume less than 1 micro 1;
- (c) maintaining the fluid sample at a predetermined temperature in the detection chamber; and
- (d) measuring an electrical parameter of an electrical circuit incorporating the detection chamber and the fluid sample in it; and
- (5) testing a food product for the presence of a predetermined type of pathogenic bacteria comprising:
 - (a) extracting a fluid sample from the food product;
- (b) feeding the extracted fluid sample to an integrated microscale biosensor;
- (c) subjecting the fluid sample to a bioseparations process to remove extraneous parties including proteins and kinds of bacteria other than the predetermined type of pathogenic bacteria;
- (d) binding bacteria of the predetermined type in the fluid sample to at least one substrate body; and
- (e) after the feeding of the extracted fluid sample to the chamber, the subjecting of the fluid sample to the bioseparations process, and the binding of the predetermined type of bacteria to the at least one substrate body, measuring an electrical parameter of an electrical circuit incorporating the detection chamber and the fluid sample to detect the presence in the fluid sample of living instances of the predetermined type of bacteria.
- USE The microscale biosensor is useful for detecting target biological substances including molecules and cells. The biosensor is useful for detecting if a microbiological substance is present in a fluid sample. The biosensor is particularly useful in methods for detecting pathogens, especially foodborne pathogens like L. monocytogenes. The biosensor is also useful for detecting or quantifying biological analytes that rely on ligand-specific binding

between a ligand and a receptor.

ADVANTAGE - Methods employing the present biosensor are able to detect pathogens in a few hours or less, possibly within minutes. The present biosensor also allows a sample of live bacteria to be distinguished from a sample of dead bacteria.

DESCRIPTION OF DRAWING(S) - The drawing shows a schematic top plan view of a biosensor.

Biochip 20

Silicon wafer substrate or body 22

Receptacles or grooves 24

Receptacles or grooves 26

Microbore tubes 28

Microbore tubes 30

Microscale channel or groove 32

Cavities or wells 34

Platinum electrodes 36

Bonding pads or electrical terminals 38

Conductors or traces 40

Glass cover 42.

Dwq.1/22

L13 ANSWER 6 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-648404 [74] WPIDS

DOC. NO. NON-CPI:

N2001-484493

DOC. NO. CPI:

C2001-191336

TITLE: Detection of analytes in a sample useful to detect chemical and biological species in air and solution

uses a three-dimensional array of a polydiacetylene backbone with a **substrate** incorporated

and monitors changes in fluorescence.

DERWENT CLASS:

A12 A89 B04 D16 S03

INVENTOR(S):

REPPY, M A; SALLER, C F; SPORN, S A

PATENT ASSIGNEE(S):

(ANAL-N) ANALYTICAL BIOLOGICAL SERVICES INC

COUNTRY COUNT: 94

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001071317 A1 20010927 (200174)* EN 54

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

AU 2001050883 A 20011003 (200210)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 200107133		WO 2001-US8790	20010320
AU 200105088		AU 2001-50883	20010320

FILING DETAILS:

PATENT NO KIND PATENT NO

AU 2001050883 A Based on

WO 200171317

PRIORITY APPLN. INFO: US 2000-190091P 20000320

2001-648404 [74] WPIDS

WO 200171317 A UPAB: 20011217 AB

NOVELTY - An analyte is detected in a sample by contacting with a three-dimensional array (e.g. liposomes) of a polydiacetylene backbone which has a substrate incorporated which has affinity for the analyte, can function as a binder to the analyte or can react with the analyte; and detecting a change in fluorescence of the array.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

(1) detecting an analyte as above but using a two-dimensional array (i.e. a film) of a polydiacetylene backbone incorporating a substrate, in which up to 90 % (optionally up to 60 %) of diacetylenes are terminated with groups specifically binding the analyte; and

(2) detecting an analyte as claimed but in which arrays are suspended in solution and analyte is detected by detecting change in polarization of the fluorescence of arrays when excited with

polarized light.

USE - The method is useful to detect chemical and biological species in air and solution e.g. small organic molecules, solvents, toxins, enzymes, peptides, bacteria and viruses etc., useful in drug discovery, medical diagnosis, food safety, pathogen detection, environmental monitoring etc. Dwq.0/5

L13 ANSWER 7 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-602793 [68] WPIDS

CROSS REFERENCE:

2002-010605 [63]

DOC. NO. NON-CPI:

N2001-449773

DOC. NO. CPI:

C2001-178619

TITLE:

Assaying a sample for a target polynucleotide or an

amplification product using an encoded bead

conjugate comprising a probe and a spectral code comprising a semiconductor nanocrystal, useful in

pharmacogenetic testing and forensics.

DERWENT CLASS:

B04 D16 L03 S03

INVENTOR(S):

BRUCHEZ, M P; LAI, J H; PHILLIPS, V E; WATSON, A R;

WONG, E Y

PATENT ASSIGNEE(S):

(QUAN-N) QUANTUM DOT CORP

COUNTRY COUNT:

94

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001071043 A1 20010927 (200168) * EN

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

AU 2001050937 A 20011003 (200210)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 20010710	43 A1	WO 2001-US9242	20010322
AU 20010509		AU 2001-50937	20010322

FILING DETAILS:

PATENT NO	KIND	•	PATENT NO
AII 20010509	37 A Based	on	WO 200171043

PRIORITY APPLN. INFO: US 2000-237000P 20000929; US 2000-191227P 20000322

AN 2001-602793 [68] WPIDS

CR 2002-010605 [63]

AB WO 200171043 A UPAB: 20020213

NOVELTY - A new method (M1) for assaying a sample for a target polynucleotide or an amplification product by contacting the sample with an encoded bead conjugate comprising a probe and a spectral code comprising a semiconductor nanocrystal. The binding between the probe and target polynucleotide results in a change in fluorescence characteristics of the bead which is measured.

DETAILED DESCRIPTION - A new method (M1) for assaying a sample for a target polynucleotide or an amplification product by contacting the sample with an encoded bead conjugate comprising a probe and a spectral code comprising a semiconductor nanocrystal. The binding between the probe and target polynucleotide results in a change in fluorescence characteristics of the bead which is measured.

In detail M1, comprises contacting the sample with an unlabelled probe polynucleotide attached to a substrate. The sample is suspected of containing the amplification product, and the amplification product comprises a first label and a capture sequence. The probe polynucleotide comprises first and second complementary regions and a third region located between the first and second complementary regions, The probe polynucleotide can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop. At least a part of the third region is complementary to at least a part of the capture sequence, and the probe polynucleotide can preferentially hybridize to the amplification product and therefore disrupt formation of the stem-loop structure under at least one set of hybridization conditions. The method then determines if the first label is associated with the substrate to determine if the amplification product is present in the sample.

INDEPENDENT CLAIMS are included for the following:

(1) an amplification product assay complex comprising a substrate comprising an unlabelled probe polynucleotide hybridized to an amplification product from a target polynucleotide, where the amplification product comprises a capture sequence and a label, where the probe polynucleotide comprises first and second complementary regions and a third region located between the first and second complementary regions, and further where the probe

polynucleotide can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop, where at least a part of the third region is hybridized to at least a part of the capture sequence, and where the stem-loop structure is not formed as a result of the probe polynucleotide being hybridized to the amplification product;

- (2) a method of forming an amplification product assay complex;
 - (3) an amplification product assay array (A1);
 - (4) a kit comprising:
- (a) a substrate attached to an unlabeled probe polynucleotide comprising first and second complementary regions and a third region located between the first and second complementary regions, where the probe polynucleotide can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop, where at least a part of the third region is complementary to at least a part of a capture sequence of an amplification product from a target polynucleotide, where the unlabeled probe polynucleotide can preferentially hybridize to the amplification product and thereby disrupt formation of the stem-loop structure under at least one set of hybridization conditions;
- (b) a reagent for incorporating a label into the amplification product;
- (c) a housing for retaining the substrate and the reagent; and
- (d) instructions provided with the housing that describe how to use the components of the kit to assay a sample for the amplification product; and
- (5) an article of manufacture, comprising a **substrate** attached to an unlabeled probe polynucleotide, where the probe comprises first and second complementary regions and a third region located between the first and second complementary regions, and the probe can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop.

USE - The methods are useful in pharmacogenetic testing, forensics, paternity testing and in screening for hereditary disorders. The methods are also useful for studying alterations of gene expression in response to a stimulus. Other applications include human population genetics, analyses of human evolutionary history, and characterization of human haplotype diversity. The methods can also be used to detect immunoglobulin class switching and hypervariable mutation of immunoglobulins, to detect polynucleotide sequences from contaminants or pathogens including bacteria, yeast and viruses, for HIV subtyping to determine the particular strains or relative amounts of particular strains infecting an individual and to detect single nucleotide polymorphisms, which may be associated with particular alleles or subsets of alleles.

The methods are also useful for mini-sequencing, and for detection mutations, including single nucleotide polymorphisms (SNPs), insertions, deletions, transitions, transversions, inversions, frame shifts, triplet repeat expansion, and chromosome rearrangements. The methods can be used to detect nucleotide sequences associated with increased risk of diseases or disorders, including cystic fibrosis, Tay-Sachs, sickle-cell anemia,

etc

ADVANTAGE - The methods are useful in multiple settings where different conjugates were used to assay for different target polynucleotides. The large number of distinguishable semiconductor nanocrystal labels allows for the simultaneous analysis of multiple labeled target polynucleotides, along with multiple different encoded bead conjugates.

The assay can be implemented in a homogenous format. This allows for higher assay throughput due to fewer manipulations of the sample and decreased cross-contamination resulting in more reliable assays and less downtime from cross-contamination.

Dwg.0/15

L13 ANSWER 8 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-616242 [71] WPIDS

CROSS REFERENCE: DOC. NO. NON-CPI:

2001-607195 [69]

DOC. NO. CPI:

N2001-459684 C2001-184468

TITLE:

New nucleic acid sensor molecule useful in diagnostic applications, nucleic acid-based

electronics and functional genomics, comprises an enzymatic nucleic acid and one or more sensors.

DERWENT CLASS: INVENTOR(S):

B04 D16 T01

BLATT, L; CHOWRIRA, B; HAEBERLI, P; MCSWIGGEN, J A;

SEIWERT, S; USMAN, N; ZINNEN, S

PATENT ASSIGNEE(S):

(RIBO-N) RIBOZYME PHARM INC

COUNTRY COUNT:

PATENT INFORMATION:

PATENT	NO	KIND	DATE	WEEK	LA	PG

95

WO 2001066721 A2 20010913 (200171)* EN 115

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW
W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ
DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE
KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO
NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ
VN YU ZA ZW

AU 2001043454 A 20010917 (200204)

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2001066721 A2	WO 2001-US7163	20010306
AU 2001043454 A	AU 2001-43454	20010306

FILING DETAILS:

PATENT NO		PATENT NO
AU 200104345		WO 200166721

PRIORITY APPLN. INFO: US 2000-187128P 20000306

AN 2001-616242 [71] WPIDS

CR 2001-607195 [69]

AB WO 200166721 A UPAB: 20020117

NOVELTY - A nucleic acid sensor molecule (I) comprising an enzymatic nucleic acid component (EC) and one or more sensor components, is new.

DETAILED DESCRIPTION - In a nucleic acid sensor molecule (I), in response to an interaction of the target signaling agent with (I), EC catalyzes a chemical reaction involving covalent attachment of at least a portion of a reporter molecule (RM) to (I), carries out a chemical reaction involving isomerization of at least a portion of RM, or catalyses a phosphorylation or dephosphorylation reaction on a non-oligonucleotide-based portion of RM.

INDEPENDENT CLAIMS are also included for:

- (1) a method involving contacting (I) and RM with a system under conditions suitable for EC to attach at least a portion of RM to (I) in the presence of a target signaling agent, to isomerize at least a portion of RM in the presence of target signaling agent, or to phosphorylate or dephosphorylate a no-oligonucleotide-based portion of RM in the presence of a target signaling agent, and assaying for the attachment of RM to (I), or assaying for the isomerization, phosphorylation or dephosphorylation reaction;
- (2) a method involving contacting (I) which comprises EC comprising a substrate binding region and a catalytic region, and a sensor component comprising a nucleic acid sequence that, upon interacting with a complementary sequence in EC, inhibits the activity of EC, and RM comprising a nucleic acid sequence complementary to the substrate binding region of EC with a system under conditions suitable for EC to catalyze cleavage of RM or to catalyze a ligation reaction involving RM in the presence of a target signaling molecule, and assaying for the cleavage and assaying for cleavage or ligation reaction;
- (3) a kit comprising (I) which comprises EC comprising a substrate binding region and a catalytic region, and a sensor component comprising a nucleic acid which inhibits the activity of EC upon interacting with a complementary sequence in EC, and RM cleavable by EC in the presence of target signaling molecule, where RM comprises a chemical moiety capable of emitting a detectable signal upon cleavage of RM;
- (4) a kit comprising (I) comprising EC including one or more sensor components, and RM, where in response to an interaction of a target signaling molecule with (I), EC catalyzes a chemical reaction involving covalent attachment of at least a portion of RM to (I), carries out a chemical reaction involving isomerization of at least a portion of RM, or catalyses a chemical reaction involving phosphorylation or dephosphorylation of a non-oligonucleotide-based portion of RM;
- (5) a method involving contacting one or more components of kit (3) or (4) with a system under conditions suitable for at least a portion of RM in (3) or (4) to be cleaved by (I) in the presence of a target molecule, or under conditions suitable for at least a portion of RM to be covalently attached to (I), isomerized by (I) or phosphorylated or dephosphorylated by (I) in the presence of a target signaling molecule;
- (6) a nucleic acid circuit comprising (I) which comprises EC and one or more sensor components, where, in response to an interaction of a target signaling agent with (I), EC catalyzes a chemical reaction involving ligation or cleavage of at least a portion of a nucleic acid based-component;
 - (7) a nucleic acid computer comprising a nucleic acid

based-component;

- (8) a method involving contacting a nucleic acid based-component with a target signaling agent under conditions suitable for (I) to ligate or cleave at least a portion of a nucleic acid based-component, and assaying the ligation or cleavage; and
- (9) isolation of (I) involving contacting a random pool of nucleic acids with a target signaling molecule and a reporter molecule, and selecting for (I) that can catalyze a chemical reaction involving covalent attachment of at least a portion of RM to (I), ligation of at least a portion of RM to (I), or phosphorylation/dephosphorylation of a non-oligonucleotide-based portion of RM by (I), in the presence of the target signaling molecule.

USE - The computer is useful for detecting a target signaling agent or to provide desired output (claimed). (I) is useful in diagnostic applications to identify the presence of genes and/or gene products which are indicative of a particular genotype and/or phenotype, for e.g. a disease state, infection, or related condition within patients, and for diagnosis of disease states or physiological abnormalities related to the expression of viral, bacterial or cellular RNA and DNA. (I) is useful in nucleic acid-based electronics, including nucleic acid-based circuits and computers, as molecular switches, and as molecular sensors capable of modulating the activity, function or physical properties of other molecules. (I) is useful for the detection of specific target signaling molecules such as nucleic acid molecules, proteins, peptides, antibodies, polysaccharides, lipids, sugars, metals, microbial or cellular metabolites, analytes, pharmaceuticals, and other organic and inorganic molecules. (I) is useful in assays to assess the specificity, toxicity and effectiveness of various small molecules, nucleoside analogs, or non-nucleic acid drugs, or their doses against validated targets or biochemical pathways, in assays involved in high-throughput screening, biochemical assays, including cellular assays, in vivo animal models, clinical trial management, and for mechanistic studies in human clinical studies. (I) is useful for the detection of pathogens, biochemicals, for example proteins, organic compounds, or inorganic compounds, in humans, plants, animals, or samples from it, in connection with environmental testing or detection of biohazards and in functional genomics, target validation and discovery, agriculture or diagnostics, for example the diagnosis of disease, or the prevention or treatment of human or animal disease. (I) is useful for detection and/or amplification of specific target signaling agents, and target signaling molecule in a system, and in DNA computing applications and nucleic acid-based electronics utilized in nucleic acid computing applications. Dwg.0/29

L13 ANSWER 9 OF 47 ACCESSION NUMBER: DOC. NO. NON-CPI: DOC. NO. CPI: TITLE:

L13 ANSWER 9 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD ACCESSION NUMBER: 2001-611185 [70] WPIDS

N2001-456251 C2001-182519

Detector for detecting a selected pathogen in a sample, comprises a substrate with a detection region on its surface, a blocking layer that blocks non-specific adsorption of pathogens, and a binder that binds the selected

pathogen.

DERWENT CLASS:

A89 B04 D16 J04 S03

INVENTOR(S): ABBOTT, N L; SKAIFE, J J

93

PATENT ASSIGNEE(S):

(WISC) WISCONSIN ALUMNI RES FOUND

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001061357 A2 20010823 (200170) * EN 52

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU

ZA ZW AU 2001043157 A 20010827 (200176)

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2001061357 A2	WO 2001-US4858	20010215
AU 2001043157 A	AU 2001-43157	20010215

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 20010431	57 A Based on	WO 200161357

PRIORITY APPLN. INFO: US 2000-182941P 20000216

AN 2001-611185 [70] WPIDS

AB WO 200161357 A UPAB: 20011129

NOVELTY - A detector (20) (I) for detecting the presence of a selected pathogen (II), comprises a substrate (S) with a detection region (DR) on its surface, where DR has microstructures comprising grooves that align liquid crystal material (LCM), a blocking layer on DR that does not disrupt the alignment of LCM but blocks non-specific adsorption of (II) on its surface, and a binder that binds (II), on DR.

DETAILED DESCRIPTION - A detector (20) comprises:

- (a) a substrate (21) (S) with DR (23) on its surface, DR having microstructures comprising grooves formed in it, that will align LCM in contact with it, where the width and depth of the grooves (26) are in the range of 10 micro m or less;
- (b) a blocking layer (BL) on the surface of DR that does not disrupt the alignment of LCM in contact with it, BL blocking non-specific adsorption of (II) to the surface; and
- (c) a binder (B) on the surface of DR, that specifically binds to (II).

INDEPENDENT CLAIMS are also included for the following:

(1) detecting the presence of (II) in a sample, by providing (S) having DR comprising a surface comprising microstructures including depressions of width and depth sized to align LCM in contact with it, where the depressions are of a size sufficient to be occupied by (II), and treating the surface of DR to provide a

layer on it that blocks non-specific binding of (II) to the surface and including (B) that specifically binds (II) to be detected; and

(2) a kit for use in the detection of (II) in a sample, comprising (S), BL, (B) and LCM, that will be aligned when in contact with DR in the absence of (II) bound to DR.

USE - (I) is useful for detecting the presence of a selected microscopic pathogen, e.g. a virus or bacteria, in a sample, by providing a substrate having DR comprising a surface comprising microstructures including depressions of width and depth sized to align LCM, the surface of DR treated to block non-specific binding of pathogens to the surface and having (B) that specifically binds the selected pathogen to be detected, applying a sample to be tested for the presence of the specific pathogen to the surface of DR and applying LCM to DR that will be aligned by the microstructures on the surface of the substrate in the absence of binding particles of the pathogen to the surface of the substrate, where the presence of selected pathogen in the sample will be manifested by a visually observable disordering of LCM caused by the pathogen particles bound to the substrate in the depressions (claimed).

ADVANTAGE - Microscopic pathogens are detected in a simple and efficient manner. The pathogen can be detected by personnel who have minimal training, and without requiring specialized laboratory facilities or equipment. Detection is provided with accurate readout in a manner that is faster than conventional serological tests. It is possible to screen for multiple microscopic pathogens in a single test. The method can be embodied in an addressable microarray, allowing the sample from a patient or from the environment to be simultaneously probed for a very broad spectrum of pathogenic agents. Moreover, by immobilizing antibodies to viral, rickettsial and bacterial surface proteins, it is possible to identify tissue targets and routes of entry of weaponized recombinant organisms faster than genetic analysis. The apparatus may also serve as a pre-screening front-end to more complex devices with embedded cells capable of detecting both biological and chemical agents.

DESCRIPTION OF DRAWING(S) - The figure shows the detector for detecting the selected pathogen in a sample. Detector 20

Substrate 21 Detection region 23 Ridges 25

Grooves 26 Dwg.1/19

L13 ANSWER 10 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

2001-638841 [73] WPIDS ACCESSION NUMBER:

N2001-477550 DOC. NO. NON-CPI: C2001-188898 DOC. NO. CPI:

TITLE:

Apparatus, useful for electrical

detection of bacterial cells, comprises substrate, microelectrodes, linkers,

counter-electrode, means for producing electrical

signal, detector and electrolyte solution.

Shears 308-4994 Searcher :

DERWENT CLASS:

A89 B04 D16 S03

INVENTOR(S):

CHOONG, V; LI, C; SAWYER, J R

PATENT ASSIGNEE(S):

(MOTI) MOTOROLA INC

COUNTRY COUNT:

94

PATENT INFORMATION:

PAT	rent no	KIND	DATE	WEEK	LA	PG
					- -	
WO	2001057	533 A2	20010809	(200173)*	EN	54

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001036625 A 20010814 (200173)

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2001057533 A2	WO 2001-US3412	20010201
AU 2001036625 A	AU 2001-36625	20010201

FILING DETAILS:

PATENT NO	KIND		PA	TENT 1	10
					
AU 20010366	25 A	Based o	n WO	20015	57533

PRIORITY APPLN. INFO: US 2000-495992 20000201

AN 2001-638841 [73] WPIDS AB WO 200157533 A UPAB: 20011211

NOVELTY - An apparatus (I) for electrical detection of bacterial cells in a sample mixture, comprising a supporting substrate (S), microelectrodes (M) in contact with (S), linking groups (L) and at least one counter-electrode (CE) in contact with (M), a means for producing an electrical signal at (M), a detector for detecting changes in electrical signal and electrolyte solution in contact with (M), (L) and CE, is new.

DETAILED DESCRIPTION - (I) comprising (S), one or a number of microelectrodes (M) in contact with (S), one or a number of linking groups (L) in contact with (M) and to which specific binding molecules have been immobilized, at least one counter-electrode (CE) in electrochemical contact with (M), a means for producing an electrical signal at each (M), a means for detecting changes in the electrical signal at each (M), and an electrolyte solution in contact with (M), (L) and CE, where bacterial cells in the sample mixture are detected by detecting a difference in the electrical signal at each (M) in the presence and absence of the sample mixture in contact with (M), is new.

INDEPENDENT CLAIMS are also included for the following:

- (1) electrical detection of bacterial cells in a sample mixture, comprising:
- (a) detecting an electrical signal (E1) in one or a number of microelectrodes in contact with (L) to which specific binding molecules have been immobilized;

- (b) exposing (M) to a sample mixture containing bacterial cells;
- (c) detecting an electrical signal (E2) in one or a number of microelectrodes;
 - (d) comparing E2 with E1; and
 - (e) determining whether E2 is different from E1;
- (2) electrical detection of viable bacterial cells in a sample. mixture, comprising:
- (a) exposing one or a number of microelectrodes in contact with (L) to which specific binding molecules have been immobilized to a sample mixture containing bacterial cells;
 - (b) detecting electrical signal (E1) in (M);
 - (c) killing the bacteria in the sample mixture exposed to (M);
 - (d) detecting an electrical signal (E2) in (M);
 - (e) comparing E2 with E1; and
 - (f) determining whether El is different from E2; and
- (3) electrical detection of bacterial cells of a particular bacterial species, subspecies or strain in a sample mixture, comprising:
- (a) detecting an electrical signal (E1) in one or a number of microelectrodes in contact with (L) to which specific binding molecules have been immobilized, where the specific binding molecules have a specificity for the bacterial cells of a particular bacterial species, subspecies or strain to be detected;
- (b) exposing (M) with a sample mixture containing bacterial cells of a particular species, subspecies or strain;
 - (c) detecting an electrical signal (E2) in (M);
 - (d) comparing E2 with E1; and
 - (e) determining whether E2 is different from E1.
- USE (I) is useful for electrical detection of bacterial cells in a sample mixture, electrical detection of viable bacterial cells in a sample mixture and electrical detection of bacterial cells of a particular bacterial species, subspecies or strain in a sample mixture (claimed). The method is useful for disease diagnostics, environmental studies, food safety analysis and pathogen detection.

ADVANTAGE - The method is an inexpensive, reliable and safe alternative to current bacterial detection methods

DESCRIPTION OF DRAWING(S) - The figure shows the schematic representation of the structure of a microelectrode in contact with a polyacrylamide gel linker moiety.

Glass capillary tube 1

Ultra fine platinum wire 2

Transition wire 3

Hookup wire 4

Epoxy plug 5

Soldered 6

Polyacrylamide gel material 7

1A, 1B/7

DERWENT INFORMATION LTD WPIDS COPYRIGHT 2002 L13 ANSWER 11 OF 47 2001-483277 [52] WPIDS

ACCESSION NUMBER:

2001-581655 [50]

CROSS REFERENCE: DOC. NO. NON-CPI:

N2001-357690

DOC. NO. CPI:

C2001-144968

TITLE:

Waveguide plate, useful in sensors for determining

many biological analytes, has, on the waveguide surface, a large coupling grating with very precise

coupling angle.

DERWENT CLASS:

B04 D16 J04 S03

INVENTOR(S):

DUVENECK, G; EDLINGER, J; HEINE, C; MAISENHOELDER,

B; PAWLAK, M

PATENT ASSIGNEE(S):

(ZEPT-N) ZEPTOSENS AG

COUNTRY COUNT:

PATENT INFORMATION:

WEEK LA PG PATENT NO KIND DATE

91

WO 2001055691 A2 20010802 (200152)* GE 39

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU

SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001037339 A 20010807 (200174)

APPLICATION DETAILS:

THIBMI NO T	CIND	APPLICATION	DATE
WO 2001055691		WO 2001-EP782 AU 2001-37339	20010125

FILING DETAILS:

PATENT NO	KIND			PAT	ENT	NO
	- -	_				
AU 20010373	39 A	Based	on	WO	2001	155691

PRIORITY APPLN. INFO: CH 2000-160

20000127

WPIDS 2001-483277 [52]

2001-581655 [50] CR

WO 200155691 A UPAB: 20011217 AB

NOVELTY - Waveguide plate (A) comprises a glass substrate (1) coated with a waveguide layer (2) and, on the surface carrying (2), at least one coupling grating, formed as a line grating with periodicity 150-1000 nm and extending, in parallel lines, at least 5

DETAILED DESCRIPTION - Waveguide plate (A) comprises a glass substrate (1) coated with a waveguide layer (2) and, on the surface carrying (2), at least one coupling grating, formed as a line grating with periodicity 150-1000 nm and extending, in parallel lines, at least 5 cm. The coupling angle (theta) changes by at most $0.1\ \mathrm{deg}$. /cm, along the line, and the absolute value of the deviation of theta from its rated value on the plate is not over 0.5 deq.

INDEPENDENT CLAIMS are also included for the following:

- (a) sensor platform (B) that includes (A);
- (b) arrangement (C) of sample containers, including (A) or (B) as base plate; and
- (c) method for simultaneous qualitative or quantitative determination of many analytes using (A), (B) or (C). USE - (A) are used as components of sensors for performing,

Shears 308-4994 Searcher :

simultaneously or sequentially, multiple quantitative or qualitative biological assays, e.g. for antigens, antibodies, nucleic acids, enzymes etc. in biological samples, water etc. Typical of many applications are in drug screening; combinatorial chemistry; binding studies; toxicity determinations; determination of gene/protein expression profiles; human and veterinary diagnosis; detection of pathogens and pollutants etc.

ADVANTAGE - The use of large, very precise gratings allows rapid analysis with reduced effort, especially no system adjustments have to be made between sequential measurements.

Dwq.0/3

L13 ANSWER 12 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-476108 [51] WPIDS

DOC. NO. CPI: C2001-142807

TITLE: New ERA binding domain polypeptides and

polynucleotides encoding them, useful as research reagents and materials for discovery of treatments

and diagnostics for diseases, or for genetic

immunization.

DERWENT CLASS: B04 D16

INVENTOR(S): LUPAS, A N; PEARCE, K H

PATENT ASSIGNEE(S): (SMIK) SMITHKLINE BEECHAM CORP; (SMIK) SMITHKLINE

BEECHAM PLC

COUNTRY COUNT: 21

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001053458 A2 20010726 (200151) * EN 279

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR W: JP US

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE
WO 2001053458 A2 WO 2001-US1786 20010117

PRIORITY APPLN. INFO: US 2000-176870P 20000118

AN 2001-476108 [51] WPIDS

AB WO 200153458 A UPAB: 20010910

NOVELTY - An isolated polypeptide (I) comprising a sequence that is either at least 70-95% identical to, comprising or having a fully defined sequence comprising an ERA binding domain given in the specification, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for

the following:

(1) an isolated polynucleotide comprising a sequence:

(a) encoding a polypeptide having at least 70-95% identity to an amino acid sequence comprising an ERA **binding** domain sequence fully given in the specification;

(b) encoding a polypeptide having at least 70-95% identity to a

nucleotide sequence encoding (I);

(c) having at least 70-95% identity to that of a polynucleotide encoding a polypeptide comprising an ERA binding domain sequence given in the specification;

- (d) encoding a polypeptide comprising an ERA binding domain sequence given in the specification; or
- (e) obtained by screening a library under stringent hybridization conditions with a labeled probe having the sequence of a polynucleotide encoding a polypeptide comprising an ERA binding domain sequence given in the specification;
 - (2) an antibody immunospecific for (I);
 - (3) a method for treating an individual:
- (a) in need of enhanced activity or expression of (I) by administering to the individual an agonist of (I) or providing to the individual an isolated polynucleotide by producing the polypeptide activity in vivo; or
- (b) having the need to inhibit activity or expression of (I) by administering an antagonist of (I), a nucleic acid that inhibits the expression of a nucleotide sequence encoding (I), a nucleic acid that inhibits the expression of a nucleotide sequence encoding (I), and/or a polypeptide that competes with the polypeptide for its ligand, substrate or receptor;
- (4) a process for diagnosing a disease or a susceptibility to a disease related to expression or activity of (I), in an individual by determining the presence of a mutation in the nucleotide sequence encoding the polypeptide in the genome of the individual, and/or analyzing for the presence or amount of the polypeptide expression in a sample derived from the individual;
- (5) a method for screening to identify compounds that activate and/or that inhibit the function of (I);
- (6) an agonist or an antagonist of the activity or expression of (I);
- (7) an expression system comprising the polynucleotide capable of producing (I) when present in a compatible host cell;
- (8) a host cell comprising the expression system of (7) or a cell membrane expressing (I);
- (9) a process for producing (I) by culturing the host cell of (8) under conditions for the production of (I);
- (10) a process for producing a host cell of (8) by transforming or transfecting a cell with the expression system of (7) under conditions allowing the production of (I); and
 - (11) a host cell produced by the process of (10).

ACTIVITY - Antimicrobial; antibacterial.

Experimental protocols are described but no results were given. MECHANISM OF ACTION - Vaccine; peptide therapy.

USE - The polynucleotides and polypeptides may be employed:

- (a) as research reagents and materials for discovery of treatments and diagnostics for diseases, particularly human diseases;
- (b) as immunogens to produce antibodies immunospecific for such polypeptides or polynucleotides;
 - (c) for genetic immunization;
- (d) to configure screening methods for detecting the effect of added compounds on the production of mRNA and/or polypeptide in cells;
- (e) in the structure-based design of an agonist, antagonist or inhibitor of the polypeptide to polynucleotide;
- (f) to interfere with the initial physical interaction between a pathogen and a eukaryotic host;
- (g) in the prevention of adhesion of bacteria to eukaryotic extracellular matrix proteins on in-dwelling devices or in wounds;
 - (h) to block ERA binding domain protein-mediated mammalian cell

invasion; to block bacterial adhesion between eukaryotic, preferably extracellular matrix proteins and bacterial ERA binding domain proteins that mediate tissue damage; and/or

(i) to block normal progression of pathogenesis in infections initiated other than by implantation of in-dwelling devices or by other surgical techniques.

The polypeptides may be used:

- (a) to identify membrane bound or soluble receptors;
- (b) as a target for the screening of antibacterial drugs; and
- (c) as antigens for vaccination of a host to produce antibodies against bacteria.

The polynucleotides may be used:

- (a) as hybridization probes for RNA, cDNA and genomic DNA to isolate full-length cDNAs and genomic clones encoding ERA binding domain:
- (b) to isolate cDNA and genomic clones of other genes having high sequence identity to the ERA binding domain gene;
- (c) for chromosome identification; and in the discovery and development of antibacterial compounds. Dwg.0/0

L13 ANSWER 13 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

CROSS REFERENCE:

2001-451868 [48] WPIDS 2001-061976 [07]; 2001-656926 [66]

DOC. NO. CPI: ·

C2001-136537

TITLE:

Detecting a nucleic acid useful in e.g. diagnosing genetic, bacterial or viral diseases, by contacting the nucleic acid with oligonucleotides attached to nanoparticles and having sequences complementary a

portion of the nucleic acid.

DERWENT CLASS:

B04 D16

93

INVENTOR(S):

ELGHANIAN, R; LETSINGER, R L; LI, Z; MIRKIN, C A;

MUCIC, R C; STORHOFF, J J; TATON, T A

PATENT ASSIGNEE(S):

(NANO-N) NANOSPHERE INC

COUNTRY COUNT:

PATENT INFORMATION:

PATENT	NO	KIND	DATE	WEEK	LA	PG

WO 2001051665 A2 20010719 (200148)* EN 229

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU

ZA ZW AU 2001032795 A 20010724 (200166)

APPLICATION DETAILS:

PATENT NO K	IND	AP	PLICATION	DATE
WO 2001051665 AU 2001032795	**-		2001 001130	20010112 20010112

FILING DETAILS:

Shears 308-4994 Searcher :

KIND PATENT NO

PATENT NO

AU 2001032795 A Based on

WO 200151665

PRIORITY APPLN. INFO: US 2001-760500 20010112; US 2000-176409P 20000113; US 2000-200161P 20000426; US

2000-603830 20000626

2001-451868 [48] WPIDS ΑN

2001-061976 [07]; 2001-656926 [66] CR

WO 200151665 A UPAB: 20011227 AB

NOVELTY - Detecting a nucleic acid having at least 2 portions, comprises contacting the nucleic acid with one or more types of nanoparticles having oligonucleotides attached to the nanoparticles and having sequences complementary to portions of the sequence of the nucleic acid.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) methods of detecting a nucleic acid having at least 2 portions comprising:
- (a) contacting the nucleic acid with one or more types of nanoparticles having oligonucleotides attached to the nanoparticles and having sequences complementary to portions of the sequence of the nucleic acid, under conditions allowing the hybridization of the oligonucleotides on the nanoparticles with the nucleic acid; and
- (b) observing a detectable change brought about by hybridization of the oligonucleotides on the nanoparticles with the nucleic acid;
- (2) kits comprising at least one container holding a composition containing at least 2 types of nanoparticles having oligonucleotides attached to it, where the first type has a sequence complementary to the sequence of a first portion of a nucleic acid, and the oligonucleotides on the second type of nanoparticles has a sequence complementary to the sequence of a second portion of the nucleic acid;
- (3) an aggregate probe comprising at least 2 types of nanoparticles having oligonucleotides attached to it, the nanoparticles of the aggregate probe are bound to each other as a result of the hybridization of some of the oligonucleotides attached to them, and at least one of the nanoparticles of the aggregate probe having oligonucleotides attached to it which have a hydrophobic group on the end not attached to the nanoparticles;
- (4) a kit comprising a container holding a core probe having at least 2 types of nanoparticles having oligonucleotides attached to it and the nanoparticles of the core probe is bound to each other as a result of the hybridization of some of the oligonucleotides attached to them;
- (5) a core probe comprising at least 2 types of nanoparticles having oligonucleotides attached to it;
 - (6) a substrate having nanoparticles attached to it;
- (7) a metallic or semiconductor nanoparticle having oligonucleotides attached to it which are labeled with fluorescent molecule at the end not attached to the nanoparticle;
- (8) a satellite probe comprising a particle having attached oligonucleotides, and probe oligonucleotides hybridized to the oligonucleotides attached to the nanoparticles;
 - (9) methods of nanofabrication;
- (10) nanomaterials or nanostructures composed of nanoparticles having oligonucleotides attached to it and being held by

oligonucleotide connectors;

- (11) a composition comprising at least 2 types of nanoparticles having oligonucleotides attached to it;
- (12) an assembly of containers holding nanoparticles having oligonucleotides attached to them;
- (13) a nanoparticle having multiple oligonucleotides attached to it:
- (14) a method of separating a selected nucleic acid having at least 2 portions from other nucleic acid;
- (15) methods of binding oligonucleotides to charged nanoparticles to produce stable nanoparticle-oligonucleotide conjugates;
- (16) nanoparticle-oligonucleotide conjugates which are nanoparticles having oligonucleotides attached to them, where the oligonucleotides are present on the surface of the nanoparticles at a surface density sufficient so that the conjugates are stable, and at least some of the oligonucleotides have sequences complementary to at least one portion of the nucleic acid or oligonucleotide sequence;
- (17) nanoparticles having oligonucleotides attached to them which comprises at least one type of recognition oligonucleotides having a sequence complementary to a portion of the nucleic acid sequence, and a type of diluent oligonucleotides; and
 - (18) methods of detecting a nucleic acid.
- USE The methods are useful for detecting nucleic acids, natural or synthetic, and modified or unmodified. The methods may also be applied in the diagnosis of genetic, bacterial and viral diseases, in forensics, in DNA sequencing, for paternity testing, for cell line authentication, and for monitoring gene therapy. The methods are further useful in research and analytical laboratories in DNA sequencing, in the field to detect the presence of specific pathogens, for quick identification of an infection to assist in drug prescription, and in homes and health centers for inexpensive first-line screening.

ADVANTAGE - The methods, which are based on observing color change with the naked eye, are cheap, fast, simple, robust (reagents are stable), do not require specialized or expensive equipment, and little or no instrumentation is required.

Dwg.0/46

L13 ANSWER 14 OF 47

WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-398175 [42] WPIDS

DOC. NO. CPI:

C2001-121143

TITLE:

Apparatus for performing biological reactions, comprises a substrate

containing surfaces for positioning array of

biomolecules, flexible layer and port which extends

from a second surface.

DERWENT CLASS: INVENTOR(S):

B04 D16 J04 HAWKINS, G W

PATENT ASSIGNEE(S):

(MOTI) MOTOROLA INC

COUNTRY COUNT:

94

PATENT INFORMATION:

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

AU 2001032641 A 20010625 (200162)

APPLICATION DETAILS:

PATENT NO		APPLICATION	DATE
WO 200104 AU 200103	4515 A2	WO 2000-US34145 AU 2001-32641	

FILING DETAILS:

PRIORITY APPLN. INFO: US 2000-605766 20000628; US 1999-464490 19991215

AN 2001-398175 [42] WPIDS

AB WO 200144515 A UPAB: 20010726

NOVELTY - Apparatus for performing biological reactions, comprises a substrate (11) containing surfaces (12,13). An array of biomolecules, positioned on the surface and a flexible layer (16) affixed to surface (12) by an adhesive layer (15), to create a reaction volume. A port (19) extends from surface (13) to the reaction volume.

USE - The apparatus is useful for performing a nucleic acid hybridization reaction used to identify pathogens, diagnose disease states, and forensic determination using gene sequences specific for a desired purpose.

ADVANTAGE - The apparatus enables performing of high capacity biological reactions. The apparatus comprises geometries that eliminate corners, therefore bubble formation is prevented, thus reducing frequency of non-specific binding and artifactual signals detected by the scanner. A thin flexible layer is contacted adjacent to a biochip present in the apparatus, thereby reducing or eliminating free surface reflection, internal reflection of illumination from scanner and dispersion or scattering of illuminated light, thereby optimizing the amount of incident light that illuminates array. This arrangement is also more economical than existing apparatus, as it minimizes need for highly polished low scattering surfaces or complex or expensive lenses, and eliminates problems associated with focus and depth-of-field in more complex optical detectors.

DESCRIPTION OF DRAWING(S) - The figure shows cross-sectional

view of the apparatus.
Substrate 11

Surfaces 12,13

Adhesive layer 15

Flexible layer 16

Port 19

Dwg.1/13

L13 ANSWER 15 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-408460 [43] WPIDS

DOC. NO. NON-CPI: N2001-302261 DOC. NO. CPI: C2001-123681

TITLE: Flow cell array for multi-analyte determination,

e.g. for drug research or food analysis, has base plate and attached bodies with channels between, forming flow cells with an inlet and an outlet

leading to a liquid reservoir.

DERWENT CLASS: A89 B04 C07 D13 D16 J04 S03

INVENTOR(S): ABEL, A P; BOPP, M A; DUVENECK, G L; EHRAT, M;

KRESBACH, G M; PAWLAK, M; SCHAERER-HERNANDEZ, N G;

SCHICK, E; SCHUERMANN-MADER, E

PATENT ASSIGNEE(S): (ZEPT-N) ZEPTOSENS AG

COUNTRY COUNT: 91

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001043875 Al 20010621 (200143)* GE 75

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001020094 A 20010625 (200162)

APPLICATION DETAILS:

171111111 110	KIND		PLICATION	DATE
WO 200104387 AU 200102009	5 A1	WO	2000-EP12668 2001-20094	

FILING DETAILS:

PATENT NO	KIND	PATEN	T NO
AU 200102009	4 A Based or	wo 20	0143875

PRIORITY APPLN. INFO: CH 2000-534 20000321; CH 1999-2316

19991217 2001-408460 [43] WPIDS

AN 2001-408460 [43] WPIDS AB WO 200143875 A UPAB: 20010801

NOVELTY - An arrangement of sample containers comprising a base plate (A) and an attached body (B) with channels between (A) and (B) arranged so as to form liquid-tight flow cell(s) with inlet(s) and outlet(s), in which at least one outlet from each flow cell leads to a reservoir which receives the liquid from the cell.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

(a) an analytical system for the determination of analyte(s), with an array as described above, arrangements for feeding samples or reagents to the sample containers in a locally-addressed fashion and detector(s) for detecting changes in measured parameters,

preferably optical, electrical, electrochemical or thermal quantities or a radioactive signal;

- (b) an analytical system for the determination of luminescence(s), with an array and feed system as above, light source(s) for excitation and detector(s) for the light emitted from one or more areas on the sensor platform;
- (c) a system for the determination of analyte(s), with an array and feed system as above, light source(s) for excitation and detector(s) for measuring a change in optical parameters, preferably refractive index (RI) and/or luminescence in the vicinity of the analyte(s);
- (d) production of a 1- or 2-dimensional array as above by assembling the base plate and attached bodies in such a way as to form a fluid-tight seal between adjacent grooves; and
- (e) detection of analytes in liquid samples with these arrangements and systems, in which samples and optionally other reagent liquids are fed into the sample containers and then flow out into a reservoir connected to the flow cell and forming a component of the sample container.

USE - For the determination of chemical, biochemical or biological analytes in screening processes for pharmaceutical research, combinatorial chemistry, clinical and preclinical development, real-time binding studies, kinetic parameters in affinity screening and research, DNA and RNA analysis and the determination of genomic and proteomic differences in the genome, e.g. single nucleotide polymorphism, measurement of protein-DNA interactions, determination of control mechanisms for m-RNA expression and protein (bio) synthesis, toxicity studies, determination of expression studies, especially for the determination of biological and chemical markers, e.g. mRNA, proteins, peptides or low-mol. wt. organic (messenger) substances, for the detection of antibodies, antigens, pathogens or bacteria in drug R and D, human and veterinary diagnostics, agrochemicals R and D, symptomatic and presymptomatic plant diagnostics and patient stratification in pharmaceutical product development, for therapeutic medicament selection and for the detection of pathogens, pollutants and irritants, especially salmonella, prions, viruses and bacteria, particularly in foods and the environment (claimed).

ADVANTAGE - An analytical system with a simple array of flow cells, enabling rapid and accurate multi-analyte determination with very small liquid samples of a very wide range of analyte types without evaporation and loss of accuracy.

DESCRIPTION OF DRAWING(S) - Cross-section of flow cell arrangement.
sample inlet; 1

sample inlet; 1
 sample outlet; 2
 recess (channel);
base plate; 4
reservoir; 5
body part 6
Dwg.1/5

L13 ANSWER 16 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD ACCESSION NUMBER: 2001-282093 [29] WPIDS DOC. NO. NON-CPI: N2001-201034

DOC. NO. NON-CPI: N2001-201034 DOC. NO. CPI: C2001-086026

TITLE: Detection of antibodies in samples useful e.g. to

measure antibody levels in serum to diagnose disease, determine vaccination efficiency or detect antibodies to recombinant proteins, by inhibition enzyme linked immunosorbant assay.

DERWENT CLASS: INVENTOR(S):

B04 D16 S03 ABRAMS, M A

PATENT ASSIGNEE(S):

(PHAA) PHARMACIA CORP

COUNTRY COUNT:

94

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA	PG
			 -	

WO 2001027621 A2 20010419 (200129)* EN 43

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000078255 A 20010423 (200147)

APPLICATION DETAILS:

11111111 110 111	ND		PLICATION	DATE
WO 2001027621 AU 2000078255	A2	WO	2000-US21992 2000-78255	

FILING DETAILS:

PATENT NO	KIND		PAT	ENT NO
AU 20000782	55 A Bas	sed on	WO	200127621

PRIORITY APPLN. INFO: US 1999-158090P 19991007

AN 2001-282093 [29] WPIDS

AB WO 200127621 A UPAB: 20010528

NOVELTY - Antibodies in a sample are detected using a new type of enzyme linked immunosorbant assay (ELISA) termed an inhibition enzyme linked immunosorbant assay (iELISA), in which purified labeled antigen and a test sample comprising an antibody are incubated with a surface coated with a purified second antibody, and antigen-binding inhibition is measured.

DETAILED DESCRIPTION - Detecting (M1) a first antibody in a sample comprises:

- (a) coating a binding surface/support with a purified second antibody to form an antibody-coated surface;
- (b) combining a predetermined amount of a purified labeled antigen and the test sample containing the first antibody;
 - (c) adding to the antibody-coated surface;
 - (d) incubating; and
 - (e) measuring antigen-binding inhibition.

An INDEPENDENT CLAIM is also included for a test kit for use with (M1), comprising an insoluble binding surface/support with a purified second antibody bound to it, a purified and labeled antigen specifically binding to and saturating the second antibody and optionally washing regents, incubating reagents and

label substrate.

USE - The method is useful to detect antibodies in samples, especially in serum from mammals, especially humans (claimed), useful e.g. to screen for elevated concentrations of endogenous antibodies to known pathogens to diagnose disease, to test sera from vaccinated humans/other animals to determine whether titers are sufficient to give protection against infection, or to detect endogenous antibodies against recombinant proteins (e.g. therapeutic proteins) which could have a neutralizing effect on the drug and drug target. It also enables detection of exogenous antibodies, useful e.g. to evaluate efficacy in disease treatment, and detection of antibodies other sample media e.g. tissue culture media, purification samples, biological fluids such as urine and saliva. The kits are especially useful for field detection of serum antibody levels, useful e.g. epidemiologically to determine particular species infected by a pathogen and/or rates of spread.

ADVANTAGE - Sensitivity of antibody measurements is increased relative to previous immunoassay techniques by eliminating background interference associated with binding of non-specific immunoglobulin. The method also increases specificity and reduces assay time and labor by incubating purified second antibody and labeled antigen simultaneously with test serum, therefore eliminating the secondary detection step normally required.

Dwg.0/9

L13 ANSWER 17 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD ACCESSION NUMBER: 2001-244826 [25] WPIDS

ACCESSION NUMBER: DOC. NO. CPI:

C2001-073497

TITLE:

Novel linear isothermal nucleic acid amplification of polynucleotide sequences by using single RNA/DNA

composite primer, which forms basis for

amplification of target sequence and optionally a

termination sequence.

DERWENT CLASS:

INVENTOR(S):

BO4 D16 KURN, N

INVENTOR(S).

PATENT ASSIGNEE(S):

(NUGE-N) NUGEN TECHNOLOGIES INC; (KURN-I) KURN N

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001020035 A2 20010322 (200125) * EN 115

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

US 6251639 B1 20010626 (200138)

AU 2000074835 A 20010417 (200140)

US 2001034048 A1 20011025 (200170)

APPLICATION DETAILS:

PATENT NO KIND

APPLICATION DATE

WO	2001020035	A2		· WO	2000-US25104	20000913
US	6251639	В1	Provisional	US	1999-153604P	19990913
			Provisional	US	2000-175780P	20000112
				US	2000-660877	20000913
ΑU	2000074835	Α		AU	2000-74835	20000913
US	2001034048	A1	Provisional	US	1999-153604P	19990913
			Provisional	US	2000-175780P	20000112
			Cont of	US	2000-660877	20000913
				US	2001-870433	20010529

FILING DETAILS:

PATENT NO F	KIND	PATENT NO
AU 2000074835	5 A Based on	WO 200120035
US 2001034048		US 6251639

PRIORITY APPLN. INFO: US 2000-175780P 20000112; US 1999-153604P 19990913; US 2000-660877 20000913; US 2001-870433 20010529

AN 2001-244826 [25] WPIDS AB WO 200120035 A UPAB: 20010508

NOVELTY - Amplifying (I) polynucleotide (PN) sequence complementary to target PN sequence (T) is new.

DETAILED DESCRIPTION - (I) comprises:

- (1) hybridizing a single stranded DNA template comprising the target sequence with a composite primer comprising an RNA portion and a 3' DNA portion;
- (2) optionally hybridizing a polynucleotide comprising a termination polynucleotide sequence to a region of the template which is 5' with respect to hybridization of the composite primer to the template;
 - (3) extending the composite primer with DNA polymerase;
- (4) cleaving the RNA portion of the annealed composite primer with an enzyme that cleaves RNA from an RNA/DNA hybrid such that another composite primer can hybridize to the template and repeat primer extension by strand displacement where multiple copies of the complementary sequence of the target sequence are produced.

INDEPENDENT CLAIMS are also included for the following:

- (1) amplifying (II) a target polynucleotide sequence comprising:
 - (a) Steps (1)-(4) of (I); and
- (b) hybridizing a polynucleotide comprising a propromoter and a region which hybridizes to the displaced primer extension product under conditions which allow transcription to occur by RNA polymerase, such that RNA transcripts are produced comprising sequences complementary to the displaced primer extension products, where multiple copies of the target sequence are produced;
- (2) characterizing (III) a sequence of interest in a target polynucleotide comprising conducting the **methods** (I) and (II) where the sequence of an RNA portion of the composite primer is known and where:
- (a) production of detectably fewer amplification products from the template as compared to the amount of amplification products from a reference template which comprises a region complementary to the RNA portion of the composite primer indicates that the target polynucleotide does not comprise a sequence complementary to the RNA

portion of the composite primer and is a sequence variant with respect to the sequence complementary to the RNA portion of the composite primer; or

- (b) production of detectably more amplification products form the template as compared to the amount of amplification products from a reference template which does not comprise a region which is complementary to the RNA portion of the composite primer indicates that the target polynucleotide comprises a sequence complementary to the RNA portion of the composite primer and is not a sequence variant with respect to the sequence complementary to the RNA portion of the composite primer;
 - (3) sequencing a target nucleotide sequence comprising:
 - (a) Steps (1) and (2) of (I);
- (b) extending the composite primer to a termination site with DNA polymerase and a mixture of dNTPs and dNTP analogs, such that primer extension is terminated upon incorporation of a dNTP analog;
- (c) cleaving the RNA portion of the annealed composite primer with an enzyme that cleaves RNA from an RNA/DNA hybrid such that another composite primer can hybridize to the template and repeat primer extension by strand displacement, where multiple copies of the complementary sequence of the target sequence are produced of varying lengths; and
- (d) analyzing the product of (a) through (d) to determine the sequence;
 - (4) sequencing a target nucleotide sequence comprising:
 - (a) Steps (1) (4) of (I);
- (b) hybridizing a polynucleotide comprising a propromoter and a region which hybridizes to the displaced primer extension product under conditions such that transcription occurs from the extension product by RNA polymerase, using a mixture of rNTPs and rNTP analogs, such that RNA transcripts are produced comprising sequences complementary to the displaced primer extension products, and such that transcription is terminated upon incorporation of an rNTP analog, whereby multiple copies of the target sequence are produced of varying lengths;
- (c) analyzing the product of steps (a) through (e) to determine the sequence;
 - (5) detecting a mutation in a target polynucleotide comprising:
 - (a) conducting (I) or (II); and
- (b) analyzing the amplified products for single stranded conformation, where a difference in conformation as compared to a reference single stranded polynucleotide indicates a mutation in the target polynucleotide;
- (6) producing a microarray, by conducting (I) and (II) and attaching the amplified products onto a solid **substrate** to make a microarray of the amplified products;
- (7) a composition comprising a complex of CP and template strand, a template switch oligonucleotide (TSO), a blocking sequence and/or a propromoter template oligonucleotide (PTO);
- (8) a reaction mixture comprising PN template, CP and DNA polymerase;
 - (9) a kit for amplification of (T), comprising CP; and
- (10) a system for amplifying (T) or its complement comprising CP, DNA polymerase and an enzyme which cleaves RNA from an RNA/DNA hybrid.
- USE The method is useful for isothermal amplification of a target nucleotide sequence or a sequence complementary to the target sequence. Linear isothermal amplification are useful for sequencing

of a defined nucleic acid target sequence and for detecting mutation in target nucleotide by analyzing the amplified products for single stranded conformation, where a difference in conformation as compared to a reference single stranded PN indicates a mutation in the target PN (claimed). The methods are also useful for qualitative detection of a nucleic acid sequence, quantitative determination of the amount of the target nucleic acid sequence, detection of the presence of defined sequence alterations, as needed for genotyping and detection of presence of various pathogens in a single biological sample. The amplified nucleic acid products are useful for genotyping and microarray preparation.

ADVANTAGE - The method does not require thermocycling in that amplification can be performed isothermally and facilitates automation and adaptation for high throughput amplification and/or analysis of nucleic acids. Sequencing based on the amplification methods are simplified by the ability to perform the reactions isothermally. The isothermal reaction is faster than that afforded by thermal cycling. Various target sequences and polymorphic sites in a single genomic DNA sample can be amplified simultaneously in a single reaction mixture.

Dwg.0/10

L13 ANSWER 18 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-182772 [18] WPIDS

DOC. NO. NON-CPI: N2001-130478

DOC. NO. CPI: C2001-054498

TITLE: New fucR polypeptides for use as research reagents

and materials for discovering treatments of

diseases, and for diagnosing bacterial infections,

specifically those caused by Streptococcus

pneumoniae.

DERWENT CLASS: B04 D16 S03

INVENTOR(S): CHAN, P F; HOLMES, D J; LONETTO, M A; TRAINI, C M;

ZALACAIN, M

PATENT ASSIGNEE(S): (SMIK) SMITHKLINE BEECHAM CORP; (SMIK) SMITHKLINE

BEECHAM PLC

COUNTRY COUNT: 19

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001007460 A1 20010201 (200118)* EN 36

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: JP

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE

WO 2001007460 A1 WO 2000-US19776 20000720

PRIORITY APPLN. INFO: US 1999-144988P 19990722

AN 2001-182772 [18] WPIDS

AB WO 200107460 A UPAB: 20010402

NOVELTY - An isolated polypeptide that is either 95% identical to, comprises, or has a sequence (I) of 251 amino acids, given in the specification, or is encoded by a recombinant polynucleotide having

a sequence (II) of 756 base pairs (bp) given in the specification, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) an isolated polynucleotide comprising:

- (i) a sequence encoding a polypeptide 95% identical to (I);
- (ii) a sequence 95% identical to a nucleotide sequence encoding
 (I);
 - (iii) a nucleotide sequence that has 95% identity to (II);

(iv) a nucleotide sequence encoding (I);

(v) the sequence (II);

- (vi) a sequence of at least 30 nucleotides in length obtained by screening a library under hybridization conditions with a probe having (II); or
 - (vii) a sequence which is a complement of the (i-vi);

(2) treating an individual:

- (a) in need of enhanced activity or expression of or immunological response to the polypeptide by administering an antagonist of the polypeptide; or
- (b) having need to inhibit activity or expression of the polypeptide by administering an antagonist of the polypeptide, a nucleic acid that inhibits the expression of a polynucleotide encoding polypeptide, a polypeptide that competes with the new polypeptide for its ligand, substrate or receptor, or a polypeptide that induces an immunological response to the polypeptide in the individual;
- (3) diagnosing or prognosing a disease or susceptibility to a disease related to the expression or activity of the polypeptide in an individual by:
- (a) determining the presence or absence of a mutation in the nucleotide sequence encoding the polypeptide; or
- (b) analyzing for the presence or amount of polypeptide expression in a sample from the individual;
- (4) producing the new polypeptide comprising culturing a host cell under conditions for the production of the polypeptide;
- (5) producing a host cell comprising an expression system or its membrane expressing the polypeptide by transforming or transfecting a cell with an expression system comprising a polynucleotide capable of producing the polypeptide when the expression system is present in a compatible host cell;
 - (6) a host cell or membrane expressing the polypeptide;

(7) an antibody immunospecific for the polypeptide;

- (8) screening compounds that agonize or inhibit the function of the polypeptide by:
- (a) measuring the **binding** of a candidate compound to the polypeptide (or to cells or membranes bearing the polypeptide) or to a fusion protein by means of a label directly or indirectly associated with the candidate compound, or in the presence of a labeled competitor
- (b) testing whether the candidate compound results in a signal generated by activation or inhibition of the polypeptide, using detection systems appropriate to the cells or cell membranes bearing the polypeptide;
- (c) mixing a candidate compound with a solution comprising the polypeptide to form a mixture, measuring activity of the polypeptide in the mixture, and comparing the activity of the mixture to a standard; or
 - (d) detecting the effect of a candidate compound on the

production of mRNA encoding the polypeptide and the polypeptide in cells, using for instance, an enzyme linked immunosorbant assay (ELISA); and

(9) an agonist or antagonist to the new polypeptide. ACTIVITY - Antibacterial. No biological data is given. MECHANISM OF ACTION - None given.

USE - The polypeptide and polynucleotides encoding it may be used as research reagents and materials for the discovery of treatments of and diagnostics for diseases, particularly human diseases. These may be particularly used in diagnosing a disease, particularly bacterial infections, specifically those caused by Streptococcus pneumoniae, and the stage and type of infection the pathogen has attained. The polynucleotides, polypeptide and antibodies that bind to or interact with the polypeptide may be used:

- (a) to configure screening methods for detecting the effect of added compounds on the production of mRNA and/or polypeptide in
- (b) to interfere with the initial interaction between a pathogen(s) and a eukaryotic cell or mammalian host responsible for sequelae of infection;
- (c) in the prevention of bacterial adhesion to extracellular proteins on in-dwelling devices or wounds;
- (d) to block bacterial adhesion between eukaryotic extracellular proteins and bacterial fucR proteins that mediate tissue damage; and/or
- (e) to block the normal progression of pathogenesis in infections initiated other than by the implantation of in-dwelling devices or by other surgical techniques.

The polypeptide is further used in identifying membrane bound or soluble receptors, and can serve as a target for screening of antibacterial drugs. The polynucleotides may be used in the discovery and development of antibacterial compounds, and to construct antisense sequences to control the expression of the coding sequence of interest. Dwq.0/0

L13 ANSWER 19 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-146292 [15] WPIDS 2000-181990 [16]; 2000-549271 [49] CROSS REFERENCE:

DOC. NO. NON-CPI: N2001-107026 DOC. NO. CPI: C2001-043201

TITLE: Detection of pathogens, DNA or

RNA useful e.g. to detect human

pathogens such as tuberculosis in serum by detecting both test and control materials using a column having a snare for each of the materials.

DERWENT CLASS: B04 D16 S03

INVENTOR(S): CHEN, H

(ACGT-N) ACGT MEDICO INC PATENT ASSIGNEE(S):

COUNTRY COUNT: PATENT INFORMATION:

> PG PATENT NO KIND DATE WEEK LA US 6174733 B1 20010116 (200115)*

APPLICATION DETAILS:

Shears 308-4994 Searcher :

above antibody) on an insoluble carrier, in known amts.; and (c) a standard soln. of a **substrate** for measuring the esterase activity, e.g. S-acetylthiophenyl or naphthyl acetate or nitrophenylacetate.

 \mbox{USE} - $\mbox{Cpds.}$ (I) are aids for rapid clinical analysis and diagnosis.

L21 859 S ABBOTT N?/AU
L22 24 S KAIFE J?/AU
L23 24 S L21 AND L22
L24 2 S (L21 OR L22) AND L1
L25 24 S L23 OR L24
L26 (10 DUPLICATES REMOVED)

L26 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2002 ACS

DUPLICATE 1

ACCESSION NUMBER: 2001:614223 CAPLUS

TITLE:

Method and apparatus for detection of

microscopic pathogens

INVENTOR(S):

Abbott, Nicholas L.; Skaife,

Justin J.

PATENT ASSIGNEE(S):

Wisconsin Alumni Research Foundation, USA

SOURCE:

PCT Int. Appl. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATE	ENT	NO.		KI	ND	DATE			A	PPLI	CATI	ON NO	0.	DATE		
WO 2	2001	0613	 57		- -	20010823			WO 2001-US4858			- <i>-</i> 8	20010215			
	W:													BZ,		CH,
														GD,		
														ΚZ,		
														MZ,		
														TR,		
														MD,		
		TM														
	RW:	GH,	GM,	KE,	LS,	MW,	ΜZ,	SD,	\$L,	SZ,	TZ,	ŪG,	ZW,	AT,	BE,	CH,
		CY,												NL,		
		TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,
		TG														
RITY	APP	LN.	INFO	. :				1	US 2	-000	1829	41	P	2000	0216	

PRIORITY APPLN. INFO.:

AB Detection apparatus for use in the detection of the presence of a selected pathogen in a sample are disclosed. Such apparatus includes: a substrate with a detection region on a surface thereof, the detection region having microstructures including grooves formed therein that will align liquid crystal material in contact therewith, the width and depth of the grooves being in the range of 10 .mu.m or less; a blocking layer on the surface of the detection region of the substrate that does not disrupt the alignment of liquid crystal material in contact therewith, the blocking layer blocking nonspecific adsorption of pathogens to the surface; and a binding agent on the surface of the detection region of the substrate, the binding agent specifically binding the selected pathogen.

L26 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 2

ACCESSION NUMBER: 2001:668371 CAPLUS

DOCUMENT NUMBER: 135:201077

TITLE: Quantitative characterization of

obliquely-deposited substrates of gold by atomic

force microscopy: influence of substrate topography on anchoring of liquid crystals

INVENTOR(S): Abbott, Nicholas L.; Skaife,

Justin J.

The Regents of the University of California, USA PATENT ASSIGNEE(S):

U.S., 33 pp. CODEN: USXXAM SOURCE:

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE US 6288392 B1 20010911 US 1999-233293 19990119

Scanning probe microscopy is used to quant. characterize structural AB anisotropy within obliquely deposited metal films. Whereas visual inspection of AFM images (real space or reciprocal space) reveals no obvious structural anisotropy within these gold films, by quant. anal. of the AFM profiles, subtle structural anisotropy is obsd. The quant. characterization provides a method to est. the influence of anisotropy on the orientations of supported mesogenic layers.

L26 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:582398 CAPLUS

DOCUMENT NUMBER: 135:253486

TITLE: Influence of Molecular-Level Interactions on the

Orientations of Liquid Crystals Supported on Nanostructured Surfaces Presenting Specifically

Bound Proteins

AUTHOR(S): Skaife, Justin J.; Abbott,

Nicholas L.

CORPORATE SOURCE: Department of Chemical Engineering, University

of Wisconsin, Madison, WI, 53706, USA

Langmuir (2001), 17(18), 5595-5604 SOURCE:

CODEN: LANGD5; ISSN: 0743-7463

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

We report an exptl. investigation of the role of mol.-level interactions in detg. the anchoring of liq. crystals supported on surfaces possessing nanometer-scale topog. on which Igs (IgG) are specifically bound to immobilized antigens. Mol.-level interactions are manipulated by using self-assembled monolayers (SAMs) of organosulfur compds. formed on thin films of gold that possess an anisotropic, nanometer-scale topog. (corrugation). We compare the orientational response of liq. crystal to the presence of anti-biotin IgG specifically bound to mixed SAMs formed from biotin-(CH2)2[(CH2)20]2NHCO(CH2)11SH and either CH3(CH2)6SH or CH3(CH2)7SH on the gold films. When using SAMs that contain 70% alkanethiolate, we measure the orientational (and thus optical) response of the liq. crystal to IqG to depend on whether the

alkanethiolate within the mixed SAM is CH3(CH2)6S or CH3(CH2)7S. conclude that in addn. to long-range (elastic) interactions that result from the nanometer-scale topog. of the gold film, mol.-level interactions controlled by the structure of the alkanethiolates influence the response of liq. crystal to the IgG specifically bound to these surfaces. The influence of the nanometer-scale topog. does, however, dominate the response of the liq. crystal. interactions appear to influence the lifetimes of line defects formed as nematic phases spread across these surfaces: the defects are obsd. to anneal quickly (.apprx. seconds) on SAMs contg. CH3(CH2)7S but slowly (> days) on those contg. CH3(CH2)6S. pinning of defects within the liq. crystal when using SAMs contg. CH3(CH2)6S causes these surfaces to be more sensitive to bound IgG than surfaces contg. CH3(CH2)7S.

REFERENCE COUNT:

THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

DUPLICATE 4

L26 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2002 ACS

37

ACCESSION NUMBER:

2001:582397 CAPLUS

DOCUMENT NUMBER:

135:253485

TITLE:

Influence of Nanometer-Scale Topography of

Surfaces on the Orientational Response of Liquid

Crystals to Proteins Specifically Bound to

Surface-Immobilized Receptors

AUTHOR(S):

Skaife, Justin J.; Brake, Jeffery M.;

Abbott, Nicholas L.

CORPORATE SOURCE:

Department of Chemical Engineering, University

of Wisconsin, Madison, WI, 53706, USA

SOURCE: Langmuir (2001), 17(18), 5448-5457

CODEN: LANGD5; ISSN: 0743-7463

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

We report procedures based on oblique deposition of gold that lead to the prepn. of ultrathin, semitransparent films of gold that possess systematic differences in their nanometer-scale topog. nanometer-scale topog. of these surfaces is controlled by the angle of incidence of the gold during the oblique deposition of each film. The topog. is quantified by using at. force microscopy (AFM) in terms of the azimuthal dependence of the contour length and local curvature of the surface. We use these surfaces to test our hypothesis that control of nanometer-scale topog. permits manipulation of the orientational response of liq. crystal to proteins bound to receptors immobilized on surfaces. We measure the orientational response of nematic phases of 4-cyano-4'pentylbiphenyl (5CB) to anti-biotin IgG bound to biotin-terminated self-assembled monolayers to depend strongly on the nanometer-scale topog. of the surfaces. The response of the liq. crystal correlates closely with quant. measures of the surface topog. obtained by AFM and thus demonstrates that it is possible to tune the sensitivity of nematic liq. crystals to the presence of specifically bound IgG by manipulating the nanometer-scale topog. of surfaces. The surfaces with the smallest local curvatures were found to be the most sensitive to the presence of bound IgG. We also calc. the anchoring energy of liq. crystal on the surfaces by using continuum elastic theory and the topog. obtained from the AFM images. Although the sensitivity of the liq. crystal to the bound protein increases with

decreasing anchoring energy, it is not possible to provide a complete account of the orientational behavior of the liq. crystal on these surfaces on the basis of continuum elastic theory.

REFERENCE COUNT: THERE ARE 38 CITED REFERENCES AVAILABLE 38

FOR THIS RECORD. ALL CITATIONS AVAILABLE

IN THE RE FORMAT

L26 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 5

2000:137840 CAPLUS ACCESSION NUMBER:

132:292361 DOCUMENT NUMBER:

TITLE: Quantitative interpretation of the optical textures of liquid crystals caused by specific

binding of immunoglobulins to surface-bound

antigens

Skaife, Justin J.; Abbott, AUTHOR(S):

Nicholas L.

Department of Chemical Engineering, University CORPORATE SOURCE:

of Wisconsin, Madison, WI, 53706, USA

Langmuir (2000), 16(7), 3529-3536 SOURCE:

CODEN: LANGD5; ISSN: 0743-7463

American Chemical Society PUBLISHER:

Journal DOCUMENT TYPE: LANGUAGE: English

The authors report a quant. anal. of the change in optical AB appearance of a supported film of liq. crystal that is induced by specific binding of an Ig (IgG) to a surface-bound antigen. The authors interpret the optical appearance to indicate the amt. of bound IgG and thus the concn. of IgG in soln. The procedure is a simple one to perform, requiring use of a CCD camera and a polarized white light source. The authors use the av. gray scale brightness of the optical appearance of the supported liq. crystal to construct an optical response curve as a function of the amt. of anti-biotin IgG bound to surface-immobilized biotin. The authors interpret the optical response curve using a model based on statistical binding of antibody to the surface and a cooperative response of the nematic liq. crystal to the bound antibody. Because the amt. of bound antibody is largely controlled by mass transport of the antibody to the surface and thus the concn. of IgG in soln., the optical appearance of the liq. crystal can be correlated to the concn. of IgG in soln. The authors measured changes in the gray scale brightness of the liq. crystal over 2 orders of magnitude of concn. of IgG in soln. (1-100 nM). The results also suggest that convection and geometry can be used to increase the dynamic range

and sensitivity of the liq. crystal to the concn. of IgG in soln. ENCE COUNT: 38 THERE ARE 38 CITED REFERENCES AVAILABLE REFERENCE COUNT:

FOR THIS RECORD. ALL CITATIONS AVAILABLE

IN THE RE FORMAT

L26 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2002 ACS 2000:333270 CAPLUS ACCESSION NUMBER:

TITLE: Using liquid crystals to image reactions on

surfaces with micrometer resolution.

Abbott, Nicholas; Shah, Rahul; AUTHOR(S):

Skaife, Justin

Department of Chemical Engineering, University CORPORATE SOURCE:

of Wisconsin, Madison, WI, 53706, USA

Book of Abstracts, 219th ACS National Meeting, SOURCE: San Francisco, CA, March 26-30, 2000 (2000),

Shears 308-4994 Searcher :

PHYS-155. American Chemical Society:

Washington, D. C. CODEN: 69CLAC

Conference; Meeting Abstract DOCUMENT TYPE:

English LANGUAGE:

Anisotropic interactions between thermotropic liq. crystals and the surfaces of solids typically cause liq. crystals to be "anchored" in one or more orientations near surfaces. Because the balance of intermol. forces acting between a liq. crystal and a surface can be easily perturbed by the transformation of state of a surface-immobilized species, and because the change in orientation of the liq. crystal can be imaged with micrometer-resoln. by using polarized light, liq. crystals can provide the basis of a general principle for the imaging of reactions on surfaces. This talk will describe the design of surfaces that permits the imaging of acid-base reactions and biospecific interactions between ligands and receptors on surfaces by using liq. crystals. Approaches to the patterning of reactants on surfaces will also be demonstrated, thereby leading to the formation of a liq. crystal grating that diffracts light upon conversion of reactant to product.

L26 ANSWER 7 OF 14 SCISEARCH COPYRIGHT 2002 ISI (R)

2000:755526 SCISEARCH ACCESSION NUMBER:

THE GENUINE ARTICLE: 317UW

Using liquid crystals to image reactions on surfaces TITLE:

with micrometer resolution.

AUTHOR: Abbott N (Reprint); Shah R; Skaife

UNIV WISCONSIN, DEPT CHEM ENGN, MADISON, WI 53706 CORPORATE SOURCE:

COUNTRY OF AUTHOR:

ABSTRACTS OF PAPERS OF THE AMERICAN CHEMICAL SOCIETY SOURCE:

(26 MAR 2000) Vol. 219, Part 2, pp. 155-PHYS. Publisher: AMER CHEMICAL SOC, 1155 16TH ST, NW,

WASHINGTON, DC 20036.

ISSN: 0065-7727. Conference; Journal

DOCUMENT TYPE:

English

LANGUAGE:

REFERENCE COUNT:

CAPLUS COPYRIGHT 2002 ACS L26 ANSWER 8 OF 14 2000:330500 CAPLUS ACCESSION NUMBER:

Soft lithography and nanoscale templating: TITLE:

Advances and applications.

Abbott, Nicholas L.; Nealey, Paul F.; AUTHOR(S):

Yang, Xiaomin; Teixeira, Ana; Skaife,

Justin J.; Kim, Seung-Ryeol

Department of Chemical Engineering, University CORPORATE SOURCE:

of Wisconsin, Madison, WI, 53706, USA

SOURCE:

Book of Abstracts, 219th ACS National Meeting, San Francisco, CA, March 26-30, 2000 (2000), IEC-044. American Chemical Society: Washington,

D. C.

CODEN: 69CLAC

DOCUMENT TYPE: Conference; Meeting Abstract

LANGUAGE: English AB Abstr. text not available.

L26 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 6

ACCESSION NUMBER:

1999:784329 CAPLUS

DOCUMENT NUMBER:

132:20781

TITLE:

Optical amplification of molecular interactions

using liquid crystals

INVENTOR(S):

Abbott, Nicholas L.; Skaife,

Justin J.; Gupta, Vinay K.; Dubrovsky,

Timothy B.; Shah, Rahul

PATENT ASSIGNEE(S):

The Regents of the University of California, USA

SOURCE:

PCT Int. Appl., 135 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PAT	CENT 1	NO.		KII	ND	DATE			F	APPLI	CATI	ON NO	ο.	DATE		
	WO	9963	 329		 A:	 1	1999	1209		V	7O 19	99-U	S125	40	1999	0604	
		W:	CA,	JP													
		RW:	AT,	BE,	CH,	CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,
			NL,	PT,	SE												
	US	6284	197		B.	1	2001	0904		Ţ	JS 19	98-1	2738:	2	1998	0731	
	ΕP	1084	394		A.	1	2001	0321		E	EP 19	99-9	2724:	3	1999	0604	
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙT,	LI,	LU,	NL,	SE,	MC,
	•		PT,	IE,	FI												
	US	2002	0042	16	A:	1	2002	0110		ţ	JS 20	01-8	9813	2	2001	0703	
PRIOR	RITY	APP	LN.	INFO	.:					US 1	1998-	9245	3	Α	1998	0605	
										US 1	998-	1273	82	Α	1998	0731	
										WO 1	999-	US12	540	W	1999	0604	

OTHER SOURCE(S): MARPAT 132:20781

App. is described which comprises a first substrate having a surface, the surface comprising a recognition moiety; a mesogenic layer oriented on the surface; and an interface between the mesogenic layer and a member selected from the group consisting of gases, liqs. solids and combinations thereof. A second substrate may be provided over the mesogenic moiety. The app. may be specifically configured for use for detecting an interaction between an analyte and a recognition moiety by detecting changes in the orientation of the mesogens occurring as a result of the interaction. Methods for detecting an analyte are described which entail contacting a recognition moiety for an analyte with a sample so that, when the analyte of interest is present, the contacting causes at least a portion of a plurality of mesogens proximate to the recognition moiety to detectably switch from a first orientation to a second orientation upon contacting the analyte with the recognition moiety; and detecting the second configuration. analyte may be selected from the group consisting of acids, bases, org. ions, inorg. ions, pharmaceuticals, herbicides, pesticides, chem. warfare agents, noxious gases, biomols., and combinations of these. App. for synthesizing and screening a library of compds. is also described which comprises a synthesis component, comprising a first substrate having a surface, and a self-assembled monolayer on the surface, the monolayer comprising a reactive functionality; and an anal. component, comprising: a second substrate having a surface, and a mesogenic layer between the surface of the first substrate and the surface of the second substrate. Libraries of compds. synthesized on a self-assembled monolayer are also claimed, as are low energy surfaces (surface energy 1-40 mJ/m2) with mesogenic

layers anchored on them. Methods for controlling the tilt of, and/or optical texture in a mesogenic layer anchored to, a haloorganosulfur moiety adsorbed on a substrate entail controlling the halogen content of the moiety.

REFERENCE COUNT:

THERE ARE 16 CITED REFERENCES AVAILABLE 16 FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1999:84969 CAPLUS

DOCUMENT NUMBER:

130:228128

TITLE:

Quantitative Characterization of Obliquely Deposited Substrates of Gold by Atomic Force Microscopy: Influence of Substrate Topography on

DUPLICATE 7

Anchoring of Liquid Crystals

AUTHOR(S):

Skaife, Justin J.; Abbott,

Nicholas L.

CORPORATE SOURCE:

Department of Chemical Engineering, University

of Wisconsin, Madison, WI, 53706, USA Chem. Mater. (1999), 11(3), 612-623 CODEN: CMATEX; ISSN: 0897-4756

SOURCE: PUBLISHER:

American Chemical Society

DOCUMENT TYPE:

LANGUAGE:

Journal English

We report the use of at. force microscopy (AFM) to characterize quant. the structural anisotropy within ultrathin (thickness of .apprx.10 nm) obliquely deposited films of gold and thereby calc. the influence of this anisotropy on the orientations of liq. crystals (LCs) supported on these surfaces. Whereas visual inspection of AFM images (real space or reciprocal space) reveals no obvious structural anisotropy within these gold films, a quant. anal. of the AFM profiles does show a subtle level of anisotropy on wavelengths comparable to the lateral dimensions of the gold grains (.apprx.30 nm). Our anal. reveals the root-mean-square (rms) slope of the surface topog. to be .apprx.1.degree. greater in a direction parallel to the direction of deposition of the gold as compared to the perpendicular direction. We also demonstrate the rms curvature of the grains of gold to be greatest in a direction parallel to deposition. Because the amplitude of the surface roughness (.apprx.2 nm) is small compared to its wavelength (.apprx.30 nm), the influence of the surface roughness on the orientations of supported LCs can be described through an elastic mechanism of anchoring. By combining the multimode Berreman-de Gennes model for the elastic free energy d. of a nematic LC with AFM profiles of the topog. of obliquely deposited gold films, we calc. the azimuthal anchoring energy of the supported LC to be .apprx.0.015 mJ/m2, a value that is consistent with ests. of anchoring energies obtained by fabrication of twisted nematic LC cells. The results reported in this paper provide a route to the characterization of surfaces with designed levels of anisotropy suitable for control of the anchoring This capability will, we believe, find application in studies aimed at exploring the use of LCs for amplification and transduction of events of mol. recognition (e.g., antigen-antibody) at surfaces.

REFERENCE COUNT:

THERE ARE 52 CITED REFERENCES AVAILABLE 52 FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

308-4994 Searcher : Shears

L26 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1999:211262 CAPLUS

DOCUMENT NUMBER: 130:317042

TITLE: Using liquid crystals as probes of

nanostructured organic surfaces

AUTHOR(S): Abbott, Nicholas L.; Shah, Rahul R.;

Gupta, Vinay K.; Skaife, Justin J.

CORPORATE SOURCE: Department of Chemical Engineering, University

of Wisconsin-Madison, Madison, WI, 53706, USA

SOURCE: Polym. Prepr. (Am. Chem. Soc., Div. Polym.

Chem.) (1999), 40(1), 425-426 CODEN: ACPPAY; ISSN: 0032-3934

PUBLISHER: American Chemical Society, Division of Polymer

Chemistry

DOCUMENT TYPE: Journal LANGUAGE: English

AB The authors describe two examples of the use of liq. crystals (LCs) as probes of nanostructured surfaces. First, the orientations of LCs supported on self-assembled monolayers (SAMs) formed from .omega.-functionalized alkanethiols are reported. When the SAMs are supported on films of gold deposited so as to possess a nanometer scale anisotropic roughness, LCs display a high level of sensitivity to the orientation and type of .omega.-functional group presented at the surface of the SAM. Second, the design of nanostructured org. surfaces is described such that specific binding of an antibody to a surface-bound antigen leads to a change in orientation of a supported LC. By using microcontact printing, surfaces were patterned with antigens such that a change in orientation of a supported LC leads to an output that is easily read by a variety of

optical methods.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1999:146175 CAPLUS

TITLE: Using liquid crystals as probes of nanostructured organic surfaces

AUTHOR(S): Abbott, Nicholas L.; Shah, Rahul R.; Gupta, Vinay K.; Skaife, Justin J.

CORPORATE SOURCE: Department of Chemical Engineering, University

of Wisconsin, Madison, WI, 53706, USA

SOURCE: Book of Abstracts, 217th ACS National Meeting,

Anaheim, Calif., March 21-25 (1999), POLY-139. American Chemical Society: Washington, D. C.

CODEN: 67GHA6

DOCUMENT TYPE: Conference; Meeting Abstract

LANGUAGE: English

AB We report two examples of the use of thermotropic liq. crystals as wet chem. probes of mol. and macromol. species supported on nanostructured surfaces. First, we report the orientations of liq. crystals supported on self-assembled monolayers (SAMs) formed from w-functionalized alkanethiols. When the SAMs are formed on films of gold deposited so as to possess a nanometer-scale, anisotropic roughness, liq. crystals display a high level of sensitivity to the orientation and type of w-functional group presented at the surface of the SAM. Second, we report the design of nanostructured, org. surfaces such that specific binding of an antibody to a

surface-bound antigen leads to a change in orientation of a supported liq. crystal.

L26 ANSWER 13 OF 14 SCISEARCH COPYRIGHT 2002 ISI (R)

1999:228382 SCISEARCH ACCESSION NUMBER:

THE GENUINE ARTICLE: 176JP

TITLE: Using liquid crystals as probes of nanostructured

organic surfaces.

AUTHOR: Abbott N L (Reprint); Shah R R; Gupta V K;

Skaife J J

UNIV WISCONSIN, DEPT CHEM ENGN, MADISON, WI 53706 CORPORATE SOURCE:

COUNTRY OF AUTHOR:

IISA

ABSTRACTS OF PAPERS OF THE AMERICAN CHEMICAL SOCIETY (21 MAR 1999) Vol. 217, Part 2, pp. 139-POLY. Publisher: AMER CHEMICAL SOC, 1155 16TH ST, NW,

WASHINGTON, DC 20036.

ISSN: 0065-7727.

DOCUMENT TYPE:

Conference; Journal

LANGUAGE:

SOURCE:

English

REFERENCE COUNT:

L26 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 8

1998:215668 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 129:2294

Optical amplification of ligand-receptor binding TITLE:

using liquid crystals

Gupta, Vinay K.; Skaife, Justin J.; AUTHOR(S):

Dubrovsky, Timothy B.; Abbott, Nicholas

L.

Dep. Chemical Eng. and Materials Science, Univ. CORPORATE SOURCE:

California, Davis, CA, 95616, USA

Science (Washington, D. C.) (1998), 279(5359), SOURCE:

2077-2080

CODEN: SCIEAS; ISSN: 0036-8075

American Association for the Advancement of PUBLISHER:

Science

DOCUMENT TYPE: Journal LANGUAGE: English

Lig. crystals (LCs) were used to amplify and transduce

receptor-mediated binding of proteins at surfaces into optical outputs. Spontaneously organized surfaces were designed so that protein mols., upon binding to ligands hosted on these surfaces,

triggered changes in the orientations of 1- to 20-.mu.m-thick films of supported LCs, thus corresponding to a reorientation of .apprx.105 to 106 mesogens per protein. Binding-induced changes in the intensity of light transmitted through the LC were easily seen with the naked eye and could be further amplified by using surfaces designed so that protein-ligand recognition causes twisted nematic LCs to untwist. This approach to the detection of ligand-receptor binding does not require labeling of the analyte, does not require the use of electroanal. app., provides a spatial resoln. of micrometers, and is sufficiently simple that it may find use in biochem. assays and imaging of spatially resolved chem. libraries.

FILE 'HOME' ENTERED AT 15:50:18 ON 06 MAR 2002

INVENTOR(S): PATENT ASSIGNEE(S): Moennig, Volker Fed. Rep. Ger. Ger. Offen., 10 pp.

SOURCE:

CODEN: GWXXBX

DOCUMENT TYPE:

Patent German

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 3840968	A1	19900607	DE 1988-3840968	19881205
DE 3840968	C2	19901004		

Monoclonal anti-idiotypic antibodies are described which mimic a AΒ genetically conserved epitope of an infective agent, which epitope only occurs in serotypes of this infective agent, induces antibody formation in the natural host, and comprises a part of an immunodominant antigen. The anti-idiotypic antibodies are prepd. by crosslinking or conjugation of a conventionally prepd. monoclonal antibody, using the product for immunization, and prodn. and selection of hybridomas secreting the anti-idiotypic antibodies. Kits for detection of an antibody to an infective agent are described which contain an immobilized monoclonal anti-idiotypic antibody and a labeled mono- or polyclonal antibody to the agent, a fluorogenic or chromogenic substrate (if the label is an enzyme), and a stop soln. Prodn. of monoclonal anti-idiotypic antibodies which mimic an epitope on European swine plaque virus is cited as an example.

LINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, EPLUS, JAPIO' ENTERED AT 15:43:57 ON 06 MAR 2002) 26 S L15

L17 **L18**

14 8 L17 NOT LT2

25 DUP REM L17 (1 DUPLICATE REMOVED) L19

14 S L19 AND BIND? L20

L20 ANSWER 1 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2002-114152 [15] WPIDS

DOC. NO. NON-CPI:

N2002-085150

DOC. NO. CPI:

C2002-034959

TITLE:

Analysis of polynucleotides in a sample using generic capture sequences comprises amplifying target polynucleotide, and utilizing the product to indirectly assay the sample for the polynucleotide.

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

LAI, J H; PHILLIPS, V E; WATSON, A R

PATENT ASSIGNEE(S):

(QUAN-N) QUANTUM DOT CORP

COUNTRY COUNT:

95

PATENT INFORMATION:

PATENT	ИО	KIND	DATE	WEEK	LA	PG

WO 2001083823 A1 20011108 (200215)* EN 85

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO

NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2001083823 A1	WO 2001-US13979	20010430

PRIORITY APPLN. INFO: US 2000-200635P 20000428

AN 2002-114152 [15] WPIDS

AB WO 200183823 A UPAB: 20020306

NOVELTY - Assaying, (M1) for an amplification product (AMP) from a first target polynucleotide, (TP), comprising providing a sample that is suspected of containing AMP, where the AMP is a polynucleotide comprising a first label and a capture sequence not present in the TP at the same position, is new.

DETAILED DESCRIPTION - A target polynucleotide, (TP), is amplified, where first primer has a tag sequence, the complement of which is formed on the opposite strand during amplification and is referred to as capture sequence (CS), and the opposite strand is referred to as an amplification product (AMP) has a label; probe conjugated to substrate specific to CS, is contacted with AMP to form AMP detection complex. Assaying (M1) for an AMP from a first TP, comprises providing a sample that is suspected of containing AMP, where the AMP is a polynucleotide comprising a first label and a capture sequence not present in the TP at the same position, where the AMP is formed by primer extension from a template, where the template comprises a complement to the TP and a target noncomplementary region, where the capture sequence is a complement to the target noncomplementary region, providing a substrate that is conjugated to a first capture probe, contacting the sample with the capture probe under a first set of hybridization conditions, where the capture probe can bind to the capture sequence under the first set of hybridization conditions and determining if the first label is associated with the substrate.

INDEPENDENT CLAIMS are also included for the following:

- (1) forming (M2) an AMP detection complex for assaying a sample for a first TP;
- (2) an amplification product detection complex comprising a capture probe polynucleotide hybridized to capture sequence of labeled amplification product from a TP, where the capture probe polynucleotide is conjugated to a substrate, where the capture sequence is not present in a region of the TP which is amplified and is introduced into the amplification product by copying a template polynucleotide, the template polynucleotide comprising a target noncomplementary region and a region complementary to the TP, where the target noncomplementary region was introduced into the template polynucleotide by extension of a primer hybridized to the target polynucleotide, the primer comprising the target noncomplementary region and where the capture sequence is complementary to the target noncomplementary region; and
- (3) a **kit** for assaying for an AMP from a TP comprises a **substrate** attached to a capture probe, a first primer comprising a 3' end a first target complementary region located at the 3' end of the first primer, and a first target noncomplementary

region that is not complementary to the first TP at a position 3' of a sequence to which the first target complementary region can hybridize, a second primer, label, housing for retaining the substrate, first primer, second primer, and a label and instruction provided with the housing that describe how to use the components of the kit to assay a sample by forming an AMP from the TP using the first and second primers that comprises the label and a capture sequence that is complementary to the target noncomplementary in the first primer, where the capture sequence can bind to the capture probe.

USE - (M2) is useful for forming an AMP detection complex for assaying a sample for first TP. The method further comprises determining if the first label is associated with the first substrate, where AMP is produced at a detectably higher level from at least one allele of a locus having at least two alleles and the first substrate preferably a first microsphere comprising a first spectral code is identified by decoding the spectral code which is preferably performed prior to, simultaneously or subsequent to determining if the first label from the second primer is associated with a substrate, first TP, preferably single-stranded or double-stranded DNA or RNA and a polymerase, preferably DNA polymerase having reverse transcriptase activity is used to form the first primer extension product (claimed). (M1) is useful for particular polynucleotide sequences, whether based on SNPs, conserved sequences, or other features or useful in a wide variety of different applications. The method is useful for pharmacogenetic testing, such methods ca be used in a forensic setting to identify the species or individual which was the source of a forensic specimen. Polynucleotide analysis methods can also be used in an anthropological setting. Paternity testing is another area, as is testing for compatibility, between prospective tissue or blood donors and patients in need, and in screening for heredity disorders. (M1) is also useful for studying gene expression in response to a stimulus. Other applications include human population genetics, analyses of human evolutionary history, and characterization of human haplotype diversity. The method is useful to detect immunoglobulin class switching and hypervariable mutation of immunoglobulins, to detect polynucleotide sequences from contaminants or pathogens including bacteria, yeast, viruses, for HIV subtyping to determine the particular strains or relative amounts of particular strains infecting an individual, and can be repeatedly to monitor changes in the individuals predominant HIV strains, such as the development of drug resistance or T cell tropism; and to detect single nucleotide polymorphisms, which may be associated with particular alleles or subsets of allele. Over 1.4 million different single nucleotide polymorphisms (SNPs) in the human population have been identified. The method is also useful for mini-sequencing and for detection of mutations. Any type of mutation can be detected, including without limitation SNPs, insertions, deletions, transitions; transversions, inversions, frame shifts, triplet repeat expansions, and chromosome rearrangements. The method is useful to detect nucleotide sequences associated with increased risk of diseases or disorders, including cystic fibrosis, Tay-Sachs, sickle-cell anemia, etc. The method is useful for any assay in which a sample can be interrogated regarding an amplification product from a target polynucleotide. Typical assays involve determine the presence of the amplification product in the sample or its relative amount, or the assays may quantitative or semi-quantitative. Results

from such assays can be used to determine the presence or amount of the target polynucleotide present in the sample. The above methods are particularly useful in multiplex settings where several TP are to be assayed. Dwg.0/15

L20 ANSWER 2 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2002-083189 [11] WPIDS

DOC. NO. CPI:

C2002-025293

TITLE:

Analyzing variant sites of target nucleic acids,

useful for identifying and detecting point

mutations, specifically those mutations correlated

with diseases e.g. cancer, by limited primer

extension.

DERWENT CLASS:

B04 D16

INVENTOR(S): PATENT ASSIGNEE(S): GLAZER, A N; XU, H (DNAS-N) DNA SCI INC

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA	PG

WO 2001092583 A1 20011206 (200211)* EN 54

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

APPLICATION DETAILS:

DATE APPLICATION PATENT NO KIND WO 2001-US18023 20010531 WO 2001092583 A1

20000602 PRIORITY APPLN. INFO: US 2000-586125

2002-083189 [11] WPIDS ΑN

WO 200192583 A UPAB: 20020215 AB

YU ZA ZW

NOVELTY - Analyzing (M1) a variant site of a target nucleic acid (TNA) by conducting template dependent extension reaction in presence of a mixture of labeled extendible nucleotide (nt) (LEN) and labeled non-extendible nts (LNEN), and detecting incorporation of labeled nt indicating identity of nt at variant site, since incorporated nt is complementary to nt at site of variation.

DETAILED DESCRIPTION - Analyzing (M1) a variant site of a target nucleic acid (TNA) by conducting template dependent extension reaction in presence of a mixture of labeled extendible nucleotide (nt) (LEN) and labeled non-extendible nts (LNEN), and detecting incorporation of labeled nt indicating identity of nt at variant site, since incorporated nt is complementary to nt at site of variation. (M1) comprises:

(a) conducting a template-dependent extension reaction comprising extending a primer (I) in the presence of TNA and a mixture of nucleotides comprising a LEN and LNEN being complementary to a different allelic form of TNA and optionally differentially

labeled, where (I) hybridizes to a segment of TNA such that the 3'-end of (I) hybridizes adjacent to the variant site of TNA, where if the LEN is complementary to the nucleotide occupying the variant site, (I) is extended by incorporation of the LEN, and can be extended further if one or more nucleotides downstream of the variant site are complementary to one of the nucleotides in the mixture, and if the LNEN is complementary to the nucleotide occupying variant site, (I) is extended by incorporation of the LNEN; and

(b) detecting incorporation of labeled nucleotide into the extended (I), the identity of the labeled nucleotide incorporated into (I) indicating the identity of the nucleotide at the variant site, where the identity of the incorporated nucleotide is determined from the label borne by the incorporated nucleotide and/or the size of the extended (I).

INDEPENDENT CLAIMS are also included for the following:

- (1) analyzing (M2) variant sites in one or more TNA comprising:
- (a) conducting several template-dependent extension reactions in the presence of several different primers, where the primers hybridize adjacent to different variant sites of TNAs and are differentially labeled, where extension reaction comprises contacting a sample containing the TNAs with one of the different labeled primers, and exposing the primer to a mixture of nucleotides comprising LEN and LNEN, where the extension reactions generate several different extension products, which are from different variant sites being distinguishable on the basis of the different labels borne by the extended primers; and
- (b) detecting incorporation of labeled nucleotides into the extension products as an indication of the nucleotides occupying the site of variation in TNAs, where the identity of the incorporated nucleotide is determined from the label borne by the incorporated nucleotide and/or the size of the extended primer; and
- (2) a **kit** (II) utilized in (M1) comprising LEN, LNEN, nucleotides complementary to different allelic forms of TNA, and a primer that hybridizes to a segment of TNA such that 3' end of the primer adjacent to the variant site of TNA.

USE - M1 is useful for analyzing a variant site of a target nucleic acid, and M2 is useful for analyzing variant sites in one or more TNA (claimed).

The method and (II) is useful for identification and detection of point mutations (e.g. somatic point mutation), specifically those mutations correlated with diseases such as diseases associated with SNPs which include sickle cell anemia, cystic fibrosis; autoimmune diseases; formation of oncogenes and cancer. For e.g. identifying whether a nucleic acid from a particular subject includes a wild-type allele or a mutant allele at a particular single nucleotide polymorphic (SNP) site. Further, the methods can be utilized to establish the genotype of the individual being tested (i.e., distinguish whether the individual is a reference-type homozygote, a heterozygote or a variant-type homozygote). The genotyping utility of the methods makes them useful within the context of medical diagnosis and prognosis. Since many SNPs are associated with various diseases and clinicians can utilize the results of the genotype study to assess the presence of disease, whether an individual is a carrier of disease, the likelihood that an individual will get a particular disease and the likely efficacy of various treatment alternatives.

The methods also have a variety of non-medical uses, such as

detecting pathogenic microorganisms, paternity testing and forensic analysis in which polymorphisms in specific games can be determined in, for e.g. blood or semen obtained from a crime scene to indicate whether a particular suspect was involved in the crime. In like manner, polymorphism analysis may be utilized in disputes to aid in determining whether a particular individual is the parent of a certain child. The methods can also be used to identify SNPs in non-humans, including, for e.g. other animals, plants, bacteria and viruses.

The methods are also useful for identifying point mutations in pathogens that could potentially result in altered pathogenicity or resistance to certain therapeutics; and to identify cells and strains having a desired genetic constitution for use in various biotechnology applications. The method is utilized as a diagnostic tool and a prognostic tool of a disease which is useful in formulating optimal treatment for the patient.

Dwg.0/6

L20 ANSWER 3 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-616242 [71] WPIDS

CROSS REFERENCE:

2001-607195 [69]

DOC. NO. NON-CPI:

N2001-459684

C2001-184468

DOC. NO. CPI: New nucleic acid sensor molecule useful in TITLE:

diagnostic applications, nucleic acid-based electronics and functional genomics, comprises an enzymatic nucleic acid and one or more sensors.

DERWENT CLASS:

B04 D16 T01

INVENTOR(S):

BLATT, L; CHOWRIRA, B; HAEBERLI, P; MCSWIGGEN, J A;

SEIWERT, S; USMAN, N; ZINNEN, S

PATENT ASSIGNEE(S):

COUNTRY COUNT:

(RIBO-N) RIBOZYME PHARM INC 95

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA	PG
				

WO 2001066721 A2 20010913 (200171)* EN 115

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001043454 A 20010917 (200204)

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2001066721 A2	WO 2001-US7163	20010306
AU 2001043454 A	AU 2001-43454	20010306

FILING DETAILS:

PATENT NO	KIND	PATENT NO
DII 20010434	54 A Based on	WO 200166721

Shears 308-4994 Searcher :

PRIORITY APPLN. INFO: US 2000-187128P 20000306

AN 2001-616242 [71] WPIDS

CR 2001-607195 [69]

AB WO 200166721 A UPAB: 20020117

NOVELTY - A nucleic acid sensor molecule (I) comprising an enzymatic nucleic acid component (EC) and one or more sensor components, is new.

DETAILED DESCRIPTION - In a nucleic acid sensor molecule (I), in response to an interaction of the target signaling agent with (I), EC catalyzes a chemical reaction involving covalent attachment of at least a portion of a reporter molecule (RM) to (I), carries out a chemical reaction involving isomerization of at least a portion of RM, or catalyses a phosphorylation or dephosphorylation reaction on a non-oligonucleotide-based portion of RM.

INDEPENDENT CLAIMS are also included for:

- (1) a method involving contacting (I) and RM with a system under conditions suitable for EC to attach at least a portion of RM to (I) in the presence of a target signaling agent, to isomerize at least a portion of RM in the presence of target signaling agent, or to phosphorylate or dephosphorylate a no-oligonucleotide-based portion of RM in the presence of a target signaling agent, and assaying for the attachment of RM to (I), or assaying for the isomerization, phosphorylation or dephosphorylation reaction;
- (2) a method involving contacting (I) which comprises EC comprising a substrate binding region and a catalytic region, and a sensor component comprising a nucleic acid sequence that, upon interacting with a complementary sequence in EC, inhibits the activity of EC, and RM comprising a nucleic acid sequence complementary to the substrate binding region of EC with a system under conditions suitable for EC to catalyze cleavage of RM or to catalyze a ligation reaction involving RM in the presence of a target signaling molecule, and assaying for the cleavage and assaying for cleavage or ligation reaction;
- (3) a kit comprising (I) which comprises EC comprising a substrate binding region and a catalytic region, and a sensor component comprising a nucleic acid which inhibits the activity of EC upon interacting with a complementary sequence in EC, and RM cleavable by EC in the presence of target signaling molecule, where RM comprises a chemical moiety capable of emitting a detectable signal upon cleavage of RM;
- (4) a kit comprising (I) comprising EC including one or more sensor components, and RM, where in response to an interaction of a target signaling molecule with (I), EC catalyzes a chemical reaction involving covalent attachment of at least a portion of RM to (I), carries out a chemical reaction involving isomerization of at least a portion of RM, or catalyses a chemical reaction involving phosphorylation or dephosphorylation of a non-oligonucleotide-based portion of RM;
- (5) a method involving contacting one or more components of kit (3) or (4) with a system under conditions suitable for at least a portion of RM in (3) or (4) to be cleaved by (I) in the presence of a target molecule, or under conditions suitable for at least a portion of RM to be covalently attached to (I), isomerized by (I) or phosphorylated or dephosphorylated by (I) in the presence of a target signaling molecule;
- (6) a nucleic acid circuit comprising (I) which comprises EC and one or more sensor components, where, in response to an

interaction of a target signaling agent with (I), EC catalyzes a chemical reaction involving ligation or cleavage of at least a portion of a nucleic acid based-component;

- (7) a nucleic acid computer comprising a nucleic acid based-component;
- (8) a method involving contacting a nucleic acid based-component with a target signaling agent under conditions suitable for (I) to ligate or cleave at least a portion of a nucleic acid based-component, and assaying the ligation or cleavage; and
- (9) isolation of (I) involving contacting a random pool of nucleic acids with a target signaling molecule and a reporter molecule, and selecting for (I) that can catalyze a chemical reaction involving covalent attachment of at least a portion of RM to (I), ligation of at least a portion of RM to (I), or phosphorylation/dephosphorylation of a non-oligonucleotide-based portion of RM by (I), in the presence of the target signaling molecule.

USE - The computer is useful for detecting a target signaling agent or to provide desired output (claimed). (I) is useful in diagnostic applications to identify the presence of genes and/or gene products which are indicative of a particular genotype and/or phenotype, for e.g. a disease state, infection, or related condition within patients, and for diagnosis of disease states or physiological abnormalities related to the expression of viral, bacterial or cellular RNA and DNA. (I) is useful in nucleic acid-based electronics, including nucleic acid-based circuits and computers, as molecular switches, and as molecular sensors capable of modulating the activity, function or physical properties of other molecules. (I) is useful for the detection of specific target signaling molecules such as nucleic acid molecules, proteins, peptides, antibodies, polysaccharides, lipids, sugars, metals, microbial or cellular metabolites, analytes, pharmaceuticals, and other organic and inorganic molecules. (I) is useful in assays to assess the specificity, toxicity and effectiveness of various small molecules, nucleoside analogs, or non-nucleic acid drugs, or their doses against validated targets or biochemical pathways, in assays involved in high-throughput screening, biochemical assays, including cellular assays, in vivo animal models, clinical trial management, and for mechanistic studies in human clinical studies. (I) is useful for the detection of pathogens, biochemicals, for example proteins, organic compounds, or inorganic compounds, in humans, plants, animals, or samples from it, in connection with environmental testing or detection of biohazards and in functional genomics, target validation and discovery, agriculture or diagnostics, for example the diagnosis of disease, or the prevention or treatment of human or animal disease. (I) is useful for detection and/or amplification of specific target signaling agents, and target signaling molecule in a system, and in DNA computing applications and nucleic acid-based electronics utilized in nucleic acid computing applications. Dwg.0/29

L20 ANSWER 4 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD ACCESSION NUMBER: 2001-611185 [70] WPIDS

ACCESSION NUMBER: 2001-611185 [70] DOC. NO. NON-CPI: N2001-456251 DOC. NO. CPI: C2001-182519

TITLE: Detector for detecting a

selected pathogen in a sample, comprises

a substrate with a detection region on its surface, a blocking layer that blocks non-specific adsorption of pathogens, and a binder that binds the selected

pathogen.

DERWENT CLASS:

A89 B04 D16 J04 S03

INVENTOR(S): PATENT ASSIGNEE(S): ABBOTT, N L; SKAIFE, J J

(WISC) WISCONSIN ALUMNI RES FOUND 93

COUNTRY COUNT:

PATENT INFORMATION:

PATENT	ИО	KIND	DATE	WEEK	LA	PG

WO 2001061357 A2 20010823 (200170)* EN

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW

AU 2001043157 A 20010827 (200176)

APPLICATION DETAILS:

ITITEMI NO	KIND	APPLICATION	DATE
WO 200106135		WO 2001-US4858	20010215
AU 200104315		AU 2001-43157	20010215

FILING DETAILS:

PATENT NO			TENT NO
AU 20010431			200161357

PRIORITY APPLN. INFO: US 2000-182941P 20000216

2001-611185 [70] WPIDS AN

AB WO 200161357 A UPAB: 20011129

> NOVELTY - A detector (20) (I) for detecting the presence of a selected pathogen (II), comprises a substrate (S) with a detection region (DR) on its surface, where DR has microstructures comprising grooves that align liquid crystal material (LCM), a blocking layer on DR that does not disrupt the alignment of LCM but blocks non-specific adsorption of (II) on its surface, and a binder that binds (II), on DR.

DETAILED DESCRIPTION - A detector (20) comprises:

- (a) a substrate (21) (S) with DR (23) on its surface, DR having microstructures comprising grooves formed in it, that will align LCM in contact with it, where the width and depth of the grooves (26) are in the range of 10 micro m or less;
- (b) a blocking layer (BL) on the surface of DR that does not disrupt the alignment of LCM in contact with it, BL blocking non-specific adsorption of (II) to the surface; and
- (c) a binder (B) on the surface of DR, that specifically binds to (II).

INDEPENDENT CLAIMS are also included for the following:

(1) detecting the presence of (II) in a sample, by providing

Shears 308-4994 Searcher :

(S) having DR comprising a surface comprising microstructures including depressions of width and depth sized to align LCM in contact with it, where the depressions are of a size sufficient to be occupied by (II), and treating the surface of DR to provide a layer on it that blocks non-specific binding of (II) to the surface and including (B) that specifically binds (II) to be detected; and

(2) a **kit** for use in the detection of (II) in a sample, comprising (S), BL, (B) and LCM, that will be aligned when in contact with DR in the absence of (II) bound to DR.

USE - (I) is useful for detecting the presence of a selected microscopic pathogen, e.g. a virus or bacteria, in a sample, by providing a substrate having DR comprising a surface comprising microstructures including depressions of width and depth sized to align LCM, the surface of DR treated to block non-specific binding of pathogens to the surface and having (B) that specifically binds the selected pathogen to be detected, applying a sample to be tested for the presence of the specific pathogen to the surface of DR and applying LCM to DR that will be aligned by the microstructures on the surface of the substrate in the absence of binding particles of the pathogen to the surface of the substrate, where the presence of selected pathogen in the sample will be manifested by a visually observable disordering of LCM caused by the pathogen particles bound to the substrate in the depressions (claimed).

ADVANTAGE - Microscopic pathogens are detected in a simple and efficient manner. The pathogen can be detected by personnel who have minimal training, and without requiring specialized laboratory facilities or equipment. Detection is provided with accurate readout in a manner that is faster than conventional serological tests. It is possible to screen for multiple microscopic pathogens in a single test. The method can be embodied in an addressable microarray, allowing the sample from a patient or from the environment to be simultaneously probed for a very broad spectrum of pathogenic agents. Moreover, by immobilizing antibodies to viral, rickettsial and bacterial surface proteins, it is possible to identify tissue targets and routes of entry of weaponized recombinant organisms faster than genetic analysis. The apparatus may also serve as a pre-screening front-end to more complex devices with embedded cells capable of detecting both biological and chemical agents.

DESCRIPTION OF DRAWING(S) - The figure shows the detector for detecting the selected pathogen in a sample.

Detector 20
Substrate 21
Detection region 23
Ridges 25

Grooves 26
Dwg.1/19
L20 ANSWER 5 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-602793 [68] WPIDS CROSS REFERENCE: 2002-010605 [63] DOC. NO. NON-CPI: N2001-449773 C2001-178619

TITLE:

Assaying a sample for a target polynucleotide or an amplification product using an encoded bead conjugate comprising a probe and a spectral code comprising a semiconductor nanocrystal, useful in

pharmacogenetic testing and forensics.

DERWENT CLASS:

B04 D16 L03 S03

INVENTOR(S):

BRUCHEZ, M P; LAI, J H; PHILLIPS, V E; WATSON, A R;

WONG, E Y

PATENT ASSIGNEE(S):

(QUAN-N) QUANTUM DOT CORP

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA E	PG
				

WO 2001071043 A1 20010927 (200168)* EN

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001050937 A 20011003 (200210)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 200107104	- :	WO 2001-US9242	20010322
AU 200105093		AU 2001-50937	20010322

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 20010509	37 A Based on	WO 200171043

PRIORITY APPLN. INFO: US 2000-237000P 20000929; US 2000-191227P 20000322

2001-602793 [68] WPIDS AN

2002-010605 [63] CR

AB WO 200171043 A UPAB: 20020213

NOVELTY - A new method (M1) for assaying a sample for a target polynucleotide or an amplification product by contacting the sample with an encoded bead conjugate comprising a probe and a spectral code comprising a semiconductor nanocrystal. The binding between the probe and target polynucleotide results in a change in fluorescence characteristics of the bead which is measured.

DETAILED DESCRIPTION - A new method (M1) for assaying a sample for a target polynucleotide or an amplification product by contacting the sample with an encoded bead conjugate comprising a probe and a spectral code comprising a semiconductor nanocrystal. The binding between the probe and target polynucleotide results in a change in fluorescence characteristics of the bead which is measured.

In detail M1, comprises contacting the sample with an unlabelled probe polynucleotide attached to a substrate. The sample is suspected of containing the amplification product, and

> 308-4994 Shears Searcher :

the amplification product comprises a first label and a capture sequence. The probe polynucleotide comprises first and second complementary regions and a third region located between the first and second complementary regions, The probe polynucleotide can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop. At least a part of the third region is complementary to at least a part of the capture sequence, and the probe polynucleotide can preferentially hybridize to the amplification product and therefore disrupt formation of the stem-loop structure under at least one set of hybridization conditions. The method then determines if the first label is associated with the substrate to determine if the amplification product is present in the sample.

INDEPENDENT CLAIMS are included for the following:

- (1) an amplification product assay complex comprising a substrate comprising an unlabelled probe polynucleotide hybridized to an amplification product from a target polynucleotide, where the amplification product comprises a capture sequence and a label, where the probe polynucleotide comprises first and second complementary regions and a third region located between the first and second complementary regions, and further where the probe polynucleotide can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop, where at least a part of the third region is hybridized to at least a part of the capture sequence, and where the stem-loop structure is not formed as a result of the probe polynucleotide being hybridized to the amplification product;
 - (2) a method of forming an amplification product assay complex;
 - (3) an amplification product assay array (A1);
 - (4) a kit comprising:
- (a) a substrate attached to an unlabeled probe polynucleotide comprising first and second complementary regions and a third region located between the first and second complementary regions, where the probe polynucleotide can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop, where at least a part of the third region is complementary to at least a part of a capture sequence of an amplification product from a target polynucleotide, where the unlabeled probe polynucleotide can preferentially hybridize to the amplification product and thereby disrupt formation of the stem-loop structure under at least one set of hybridization conditions;
- (b) a reagent for incorporating a label into the amplification product;
- (c) a housing for retaining the substrate and the reagent; and
- (d) instructions provided with the housing that describe how to use the components of the **kit** to assay a sample for the amplification product; and
- (5) an article of manufacture, comprising a substrate attached to an unlabeled probe polynucleotide, where the probe comprises first and second complementary regions and a third region located between the first and second complementary regions, and the probe can form a stem-loop structure in which the first and second complementary regions hybridize to each other to form a stem and the third region forms a loop.

USE - The methods are useful in pharmacogenetic testing, forensics, paternity testing and in screening for hereditary disorders. The methods are also useful for studying alterations of gene expression in response to a stimulus. Other applications include human population genetics, analyses of human evolutionary history, and characterization of human haplotype diversity. The methods can also be used to detect immunoglobulin class switching and hypervariable mutation of immunoglobulins, to detect polynucleotide sequences from contaminants or pathogens including bacteria, yeast and viruses, for HIV subtyping to determine the particular strains or relative amounts of particular strains infecting an individual and to detect single nucleotide polymorphisms, which may be associated with particular alleles or subsets of alleles.

The methods are also useful for mini-sequencing, and for detection mutations, including single nucleotide polymorphisms (SNPs), insertions, deletions, transitions, transversions, inversions, frame shifts, triplet repeat expansion, and chromosome rearrangements. The methods can be used to detect nucleotide sequences associated with increased risk of diseases or disorders, including cystic fibrosis, Tay-Sachs, sickle-cell anemia, etc.

ADVANTAGE - The methods are useful in multiple settings where different conjugates were used to assay for different target polynucleotides. The large number of distinguishable semiconductor nanocrystal labels allows for the simultaneous analysis of multiple labeled target polynucleotides, along with multiple different encoded bead conjugates.

The assay can be implemented in a homogenous format. This allows for higher assay throughput due to fewer manipulations of the sample and decreased cross-contamination resulting in more reliable assays and less downtime from cross-contamination.

Dwg.0/15

L20 ANSWER 6 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-451868 [48] WPIDS

CROSS REFERENCE:

2001-061976 [07]; 2001-656926 [66]

DOC. NO. CPI:

C2001-136537

TITLE:

Detecting a nucleic acid useful in e.g. diagnosing genetic, bacterial or viral diseases, by contacting the nucleic acid with oligonucleotides attached to nanoparticles and having sequences complementary a

portion of the nucleic acid.

DERWENT CLASS:

B04 D16

INVENTOR(S):

ELGHANIAN, R; LETSINGER, R L; LI, Z; MIRKIN, C A;

MUCIC, R C; STORHOFF, J J; TATON, T A

PATENT ASSIGNEE(S):

(NANO-N) NANOSPHERE INC

COUNTRY COUNT:

93

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001051665 A2 20010719 (200148)* EN 229

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU

ZA ZW AU 2001032795 A 20010724 (200166)

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2001051665 A2	WO 2001-US1190	20010112
AU 2001032795 A	AU 2001-32795	20010112

FILING DETAILS:

	KIND			ON TN
AU 20010327			•	00151665

PRIORITY APPLN. INFO: US 2001-760500 20010112; US 2000-176409P 20000113; US 2000-200161P 20000426; US

2000-603830 20000626

AN 2001-451868 [48] WPIDS

CR 2001-061976 [07]; 2001-656926 [66]

AB WO 200151665 A UPAB: 20011227

NOVELTY - Detecting a nucleic acid having at least 2 portions, comprises contacting the nucleic acid with one or more types of nanoparticles having oligonucleotides attached to the nanoparticles and having sequences complementary to portions of the sequence of the nucleic acid.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) methods of detecting a nucleic acid having at least 2 portions comprising:
- (a) contacting the nucleic acid with one or more types of nanoparticles having oligonucleotides attached to the nanoparticles and having sequences complementary to portions of the sequence of the nucleic acid, under conditions allowing the hybridization of the oligonucleotides on the nanoparticles with the nucleic acid; and
- (b) observing a detectable change brought about by hybridization of the oligonucleotides on the nanoparticles with the nucleic acid;
- (2) kits comprising at least one container holding a composition containing at least 2 types of nanoparticles having oligonucleotides attached to it, where the first type has a sequence complementary to the sequence of a first portion of a nucleic acid, and the oligonucleotides on the second type of nanoparticles has a sequence complementary to the sequence of a second portion of the nucleic acid;
- (3) an aggregate probe comprising at least 2 types of nanoparticles having oligonucleotides attached to it, the nanoparticles of the aggregate probe are bound to each other as a result of the hybridization of some of the oligonucleotides attached to them, and at least one of the nanoparticles of the aggregate probe having oligonucleotides attached to it which have a hydrophobic group on the end not attached to the nanoparticles;
- (4) a kit comprising a container holding a core probe having at least 2 types of nanoparticles having oligonucleotides attached to it and the nanoparticles of the core probe is bound to each other as a result of the hybridization of some of the oligonucleotides attached to them;

- (5) a core probe comprising at least 2 types of nanoparticles having oligonucleotides attached to it;
 - (6) a substrate having nanoparticles attached to it;
- (7) a metallic or semiconductor nanoparticle having oligonucleotides attached to it which are labeled with fluorescent molecule at the end not attached to the nanoparticle;
- (8) a satellite probe comprising a particle having attached oligonucleotides, and probe oligonucleotides hybridized to the oligonucleotides attached to the nanoparticles;
 - (9) methods of nanofabrication;
- (10) nanomaterials or nanostructures composed of nanoparticles having oligonucleotides attached to it and being held by oligonucleotide connectors;
- (11) a composition comprising at least 2 types of nanoparticles having oligonucleotides attached to it;
- (12) an assembly of containers holding nanoparticles having oligonucleotides attached to them;
- (13) a nanoparticle having multiple oligonucleotides attached to it:
- (14) a method of separating a selected nucleic acid having at least 2 portions from other nucleic acid;
- (15) methods of **binding** oligonucleotides to charged nanoparticles to produce stable nanoparticle-oligonucleotide conjugates;
- (16) nanoparticle-oligonucleotide conjugates which are nanoparticles having oligonucleotides attached to them, where the oligonucleotides are present on the surface of the nanoparticles at a surface density sufficient so that the conjugates are stable, and at least some of the oligonucleotides have sequences complementary to at least one portion of the nucleic acid or oligonucleotide sequence;
- (17) nanoparticles having oligonucleotides attached to them which comprises at least one type of recognition oligonucleotides having a sequence complementary to a portion of the nucleic acid sequence, and a type of diluent oligonucleotides; and
 - (18) methods of detecting a nucleic acid.
- USE The methods are useful for detecting nucleic acids, natural or synthetic, and modified or unmodified. The methods may also be applied in the diagnosis of genetic, bacterial and viral diseases, in forensics, in DNA sequencing, for paternity testing, for cell line authentication, and for monitoring gene therapy. The methods are further useful in research and analytical laboratories in DNA sequencing, in the field to detect the presence of specific pathogens, for quick identification of an infection to assist in drug prescription, and in homes and health centers for inexpensive first-line screening.

ADVANTAGE - The methods, which are based on observing color change with the naked eye, are cheap, fast, simple, robust (reagents are stable), do not require specialized or expensive equipment, and little or no instrumentation is required.

Dwg.0/46

L20 ANSWER 7 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-282093 [29] WPIDS

DOC. NO. NON-CPI: N2001-201034 DOC. NO. CPI: C2001-086026

TITLE: Detection of antibodies in samples useful e.g. to measure antibody levels in serum to diagnose

disease, determine vaccination efficiency or detect antibodies to recombinant proteins, by inhibition enzyme linked immunosorbant assay.

DERWENT CLASS: INVENTOR(S):

B04 D16 S03 ABRAMS, M A

PATENT ASSIGNEE(S):

(PHAA) PHARMACIA CORP

COUNTRY COUNT:

94

PATENT INFORMATION:

PATENT	NO	KIND	DATE	WEEK	LA	PG

WO 2001027621 A2 20010419 (200129)* EN 43

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000078255 A 20010423 (200147)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 20010276 AU 20000782	21 A2	WO 2000-US21992 AU 2000-78255	

FILING DETAILS:

PATENT NO		PATENT NO
AU 20000782		WO 200127621

PRIORITY APPLN. INFO: US 1999-158090P 19991007

AN 2001-282093 [29] WPIDS

AB WO 200127621 A UPAB: 20010528

NOVELTY - Antibodies in a sample are detected using a new type of enzyme linked immunosorbant assay (ELISA) termed an inhibition enzyme linked immunosorbant assay (iELISA), in which purified labeled antigen and a test sample comprising an antibody are incubated with a surface coated with a purified second antibody, and antigen-binding inhibition is measured.

DETAILED DESCRIPTION - Detecting (M1) a first antibody in a sample comprises:

- (a) coating a binding surface/support with a purified second antibody to form an antibody-coated surface;
- (b) combining a predetermined amount of a purified labeled antigen and the test sample containing the first antibody;
 - (c) adding to the antibody-coated surface;
 - (d) incubating; and
 - (e) measuring antigen-binding inhibition.

An INDEPENDENT CLAIM is also included for a test kit for use with (M1), comprising an insoluble binding surface/support with a purified second antibody bound to it, a purified and labeled antigen specifically binding to and saturating the second antibody and optionally washing regents, incubating reagents and label substrate.

USE - The method is useful to detect antibodies in samples, especially in serum from mammals, especially humans (claimed), useful e.g. to screen for elevated concentrations of endogenous antibodies to known pathogens to diagnose disease, to test sera from vaccinated humans/other animals to determine whether titers are sufficient to give protection against infection, or to detect endogenous antibodies against recombinant proteins (e.g. therapeutic proteins) which could have a neutralizing effect on the drug and drug target. It also enables detection of exogenous antibodies, useful e.g. to evaluate efficacy in disease treatment, and detection of antibodies other sample media e.g. tissue culture media, purification samples, biological fluids such as urine and saliva. The kits are especially useful for field detection of serum antibody levels, useful e.g. epidemiologically to determine particular species infected by a pathogen and/or rates of spread.

ADVANTAGE - Sensitivity of antibody measurements is increased relative to previous immunoassay techniques by eliminating background interference associated with **binding** of non-specific immunoglobulin. The method also increases specificity and reduces assay time and labor by incubating purified second antibody and labeled antigen simultaneously with test serum, therefore eliminating the secondary detection step normally required.

Dwg.0/9

L20 ANSWER 8 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-248306 [26] WPIDS

DOC. NO. NON-CPI:

N2001-176898

DOC. NO. CPI:

C2001-075067

TITLE:

Detecting test substances useful for

detecting, e.g. pathogens,

comprises adding test liquid, enzyme-labeled

specific binding substance and

substrate liquid to their respective

addition parts on immunological inspection piece.

DERWENT CLASS: B04 D16 S03

PATENT ASSIGNEE(S):

(NITL) NITTO DENKO CORP

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

JP 2001013140 A 20010119 (200126)* 8

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE

JP 2001013140 A JP 1999-182412 19990628

PRIORITY APPLN. INFO: JP 1999-182412 19990628

AN 2001-248306 [26] WPIDS

AB JP2001013140 A UPAB: 20010515

NOVELTY - Detecting test substance (T) comprises using base material having fixed part (I) comprising (T) coupled to first immunological reagent (R1), substrate (S) addition part (AP) (II),

enzyme(E)-labeled specific **binding** substance (III) which couples (R2) and (E), to (T). Test liquid-(III) AP is arranged between (I) and (II). The test liquid, (III), (S)-liquid are added in this order in their respective APs.

DETAILED DESCRIPTION - Detecting (T) involves use of immunological inspection piece which has water absorptive base material on which is present, a fixed part (I) comprising (T) to which a first immunological reagent (R1) is coupled, substrate (S) liquid addition part (II) for adding (S), an enzyme(E)-labeled specific binding substance (III) which couples a second immunological reagent (R2) and enzyme, to (T). The addition part of (III) (1) is arranged between (I) and (II).

An INDEPENDENT CLAIM is also included for a **kit** for detecting substances.

USE - For detecting test substances such as pathogens.

ADVANTAGE - The method is rapid and simple. Dwg.0/2

L20 ANSWER 9 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-244826 [25] WPIDS

DOC. NO. CPI:

C2001-073497

TITLE:

Novel linear isothermal nucleic acid amplification of polynucleotide sequences by using single RNA/DNA composite primer, which forms basis for

composite primer, which forms basis for amplification of target sequence and optionally a

termination sequence.

DERWENT CLASS:

B04 D16

INVENTOR(S):

KURN, N

94

PATENT ASSIGNEE(S):

(NUGE-N) NUGEN TECHNOLOGIES INC; (KURN-I) KURN N

COUNTRY COUNT:

__ _ _ _

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001020035 A2 20010322 (200125) * EN 115

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN

YU ZA ZW

US 6251639 B1 20010626 (200138)

AU 2000074835 A 20010417 (200140)

US 2001034048 A1 20011025 (200170)

APPLICATION DETAILS:

PAS	TENT NO K	IND		AP	PLICATION	DATE
WO	2001020035	A2		WO	2000-US25104	20000913
US	6251639	В1	Provisional	US	1999-153604P	19990913
			Provisional	US	2000-175780P	20000112
				US	2000-660877	20000913
ΑU	2000074835	Α		Αŭ	2000-74835	20000913
US	2001034048	A1	Provisional	US	1999-153604P	19990913
			Provisional	US	2000-175780P	20000112

Cont of

US 2000-660877 20000913 US 2001-870433 20010529

FILING DETAILS:

PATENT NO KIND PATENT NO

AU 2000074835 A Based on WO 200120035
US 2001034048 A1 Cont of US 6251639

PRIORITY APPLN. INFO: US 2000-175780P 20000112; US 1999-153604P 19990913; US 2000-660877 20000913; US

2001-870433 20010529

AN 2001-244826 [25] WPIDS

AB WO 200120035 A UPAB: 20010508

NOVELTY - Amplifying (I) polynucleotide (PN) sequence complementary to target PN sequence (T) is new.

DETAILED DESCRIPTION - (I) comprises:

- (1) hybridizing a single stranded DNA template comprising the target sequence with a composite primer comprising an RNA portion and a 3' DNA portion;
- (2) optionally hybridizing a polynucleotide comprising a termination polynucleotide sequence to a region of the template which is 5' with respect to hybridization of the composite primer to the template;
 - (3) extending the composite primer with DNA polymerase;
- (4) cleaving the RNA portion of the annealed composite primer with an enzyme that cleaves RNA from an RNA/DNA hybrid such that another composite primer can hybridize to the template and repeat primer extension by strand displacement where multiple copies of the complementary sequence of the target sequence are produced.

INDEPENDENT CLAIMS are also included for the following:

- (1) amplifying (II) a target polynucleotide sequence comprising:
 - (a) Steps (1)-(4) of (I); and
- (b) hybridizing a polynucleotide comprising a propromoter and a region which hybridizes to the displaced primer extension product under conditions which allow transcription to occur by RNA polymerase, such that RNA transcripts are produced comprising sequences complementary to the displaced primer extension products, where multiple copies of the target sequence are produced;
- (2) characterizing (III) a sequence of interest in a target polynucleotide comprising conducting the methods (I) and (II) where the sequence of an RNA portion of the composite primer is known and where:
- (a) production of detectably fewer amplification products from the template as compared to the amount of amplification products from a reference template which comprises a region complementary to the RNA portion of the composite primer indicates that the target polynucleotide does not comprise a sequence complementary to the RNA portion of the composite primer and is a sequence variant with respect to the sequence complementary to the RNA portion of the composite primer; or
- (b) production of detectably more amplification products form the template as compared to the amount of amplification products from a reference template which does not comprise a region which is complementary to the RNA portion of the composite primer indicates that the target polynucleotide comprises a sequence complementary to

the RNA portion of the composite primer and is not a sequence variant with respect to the sequence complementary to the RNA portion of the composite primer;

- (3) sequencing a target nucleotide sequence comprising:
- (a) Steps (1) and (2) of (I);
- (b) extending the composite primer to a termination site with DNA polymerase and a mixture of dNTPs and dNTP analogs, such that primer extension is terminated upon incorporation of a dNTP analog;
- (c) cleaving the RNA portion of the annealed composite primer with an enzyme that cleaves RNA from an RNA/DNA hybrid such that another composite primer can hybridize to the template and repeat primer extension by strand displacement, where multiple copies of the complementary sequence of the target sequence are produced of varying lengths; and
- (d) analyzing the product of (a) through (d) to determine the sequence;
 - (4) sequencing a target nucleotide sequence comprising:
 - (a) Steps (1)-(4) of (I);
- (b) hybridizing a polynucleotide comprising a propromoter and a region which hybridizes to the displaced primer extension product under conditions such that transcription occurs from the extension product by RNA polymerase, using a mixture of rNTPs and rNTP analogs, such that RNA transcripts are produced comprising sequences complementary to the displaced primer extension products, and such that transcription is terminated upon incorporation of an rNTP analog, whereby multiple copies of the target sequence are produced of varying lengths;
- (c) analyzing the product of steps (a) through (e) to determine the sequence;
 - (5) detecting a mutation in a target polynucleotide comprising:
 - (a) conducting (I) or (II); and
- (b) analyzing the amplified products for single stranded conformation, where a difference in conformation as compared to a reference single stranded polynucleotide indicates a mutation in the target polynucleotide;
- (6) producing a microarray, by conducting (I) and (II) and attaching the amplified products onto a solid **substrate** to make a microarray of the amplified products;
- (7) a composition comprising a complex of CP and template strand, a template switch oligonucleotide (TSO), a blocking sequence and/or a propromoter template oligonucleotide (PTO);
- (8) a reaction mixture comprising PN template, CP and DNA polymerase;
- (9) a **kit** for amplification of (T), comprising CP;
- (10) a system for amplifying (T) or its complement comprising CP, DNA polymerase and an enzyme which cleaves RNA from an RNA/DNA hybrid.
- USE The method is useful for isothermal amplification of a target nucleotide sequence or a sequence complementary to the target sequence. Linear isothermal amplification are useful for sequencing of a defined nucleic acid target sequence and for detecting mutation in target nucleotide by analyzing the amplified products for single stranded conformation, where a difference in conformation as compared to a reference single stranded PN indicates a mutation in the target PN (claimed). The methods are also useful for qualitative detection of a nucleic acid sequence, quantitative determination of the amount of the target nucleic acid sequence, detection of the

presence of defined sequence alterations, as needed for genotyping and detection of presence of various pathogens in a single biological sample. The amplified nucleic acid products are useful for genotyping and microarray preparation.

ADVANTAGE - The method does not require thermocycling in that amplification can be performed isothermally and facilitates automation and adaptation for high throughput amplification and/or analysis of nucleic acids. Sequencing based on the amplification methods are simplified by the ability to perform the reactions isothermally. The isothermal reaction is faster than that afforded by thermal cycling. Various target sequences and polymorphic sites in a single genomic DNA sample can be amplified simultaneously in a single reaction mixture. Dwg.0/10

L20 ANSWER 10 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

WPIDS 2001-146292 [15]

CROSS REFERENCE:

2000-181990 [16]; 2000-549271 [49]

DOC. NO. NON-CPI: DOC. NO. CPI:

N2001-107026 C2001-043201

TITLE:

Detection of pathogens, DNA or RNA useful e.g. to detect human

pathogens such as tuberculosis in serum by

detecting both test and control materials using a column having a snare for each of the materials.

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

CHEN, H

PATENT ASSIGNEE(S):

(ACGT-N) ACGT MEDICO INC

COUNTRY COUNT:

PATENT INFORMATION:

KIND DATE PATENT NO

PG WEEK B1 20010116 (200115)*

APPLICATION DETAILS:

US 6174733

PATENT	NO I	KIND	APE	PLICATION	DATE
US 6174	1733	B1	US	1998-93532	19980608

PRIORITY APPLN. INFO: GB 1997-11941 19970609

AN 2001-146292 [15] WPIDS

2000-181990 [16]; 2000-549271 [49] CR

6174733 B UPAB: 20010317 AB

NOVELTY - A detection method for detecting the presence of at least two predetermined materials comprises detecting two predetermined materials, one of which is a control, using a column which has a snare for each of the materials.

DETAILED DESCRIPTION - The method comprises:

- (a) introducing a test sample comprising at least one control material into a column having a snare for each predetermined material, the snare having a capture material which binds specifically with the associated predetermined material;
 - (b) washing the test column to remove unbound materials; and
- (c) detecting bound materials on each of the snares, optionally by adding a label for each of the bound materials to form labeled

308-4994 Shears Searcher :

bound materials and detecting labeled bound materials.

INDEPENDENT CLAIM are also included for:

- (1) a column for analyzing at least one pathogen by the method, comprising at least two spatially separated snares, one having a control capture material and the other a pathogen capture material to enable detection of control material and a pathogen of interest; and
 - (2) a kit which comprises:
- (a) a column for analysis of at least one pathogen in which the column has at least two snares. and the snares are separated spatially one from another so that the snares are not in contact with one another, on of the snares having on it a first control capture material for detecting the presence of a first control material, and the other of the snares having on them a pathogen capture material for detecting a pathogen; and
- (b) reagents for detecting the presence of materials selected from:
- (i) reagents for detecting the presence of the control pathogen and the test pathogen; and
- (ii) reagents for detecting the presence of the first control capture material and the pathogen capture material after the first control capture material and the pathogen capture material have been bound and then unbound from the first control material and the pathogen material.

USE - The method is useful to detect
pathogens e.g. human pathogens such as
tuberculosis, especially by detecting proteins, DNA or RNA
in serum. For example, a DNA method may be used to diagnose herpes
simplex virus and an RNA method to diagnose HIV. The method may also
be useful in veterinary medicine and to detect chemicals such as
drugs, carcinogens, pollutants etc.

ADVANTAGE - Unlike previous pathogen detection techniques, the method is rapid and is sensitive to any error in the method because of the inclusion of a control snare.

Dwg.0/11

L20 ANSWER 11 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1998-568229 [48] WPIDS

DOC. NO. NON-CPI: N1998-442113
DOC. NO. CPI: C1998-170698

TITLE: Assay for e.g. pathogen(s), toxins, lymphocytes or cancer cells - using polyvinylidene di fluoride

surface to which lipid based receptor is attached.

DERWENT CLASS: A96 B04 D16 S03 INVENTOR(S): CHATTERJEE, S

PATENT ASSIGNEE(S): (UYJO) UNIV JOHNS HOPKINS

COUNTRY COUNT: 80

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 9835233 A1 19980813 (199848) * EN 33

RW: AT BE CH DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE GH GM GW HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT

LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG UZ VN YU ZW AU 9862627 A 19980826 (199902)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9835233 AU 9862627	A1	WO 1998-US1977	19980206 19980206

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9862627	A Based on	WO 9835233

PRIORITY APPLN. INFO: US 1997-37553P 19970211; US 1997-38145P 19970210

AN 1998-568229 [48] WPIDS

AB WO 9835233 A UPAB: 19981210

Assaying a test sample for the presence of a pathogen comprises: (a) applying a glycosphingolipid (GL) which is a receptor for a characteristic component (CC) of the pathogen and which specifically binds the (CC) to a polyvinylidene difluoride (PVDF) surface; (b) applying a liquid reaction medium comprising the test sample to the surface; (c) removing excess liquid medium from the surface; (d) incubating the surface with an antibody which specifically binds to the CC and (e) detecting the presence of the antibody on the surface which indicates the presence of CC in the test sample. Also claimed is a kit for detecting a pathogen which comprises a PVDF surface, a purified sample of a GL which specifically binds to a characteristic component of the pathogen and an antibody which specifically binds to the characteristic component of the pathogen.

The characteristic component is preferably a bacterial toxin (BT). The BT is preferably Vibrio cholera and the GL is GM1 ganglioside, the BT is Staphylococcal enterotoxin-B and the GL is digalactosylceramide, the BT is Staphylococcal enterotoxin-A and the GL is globotriosylamide, the BT is Verocytotoxin-2 and the GL is globotriosylceramide, the BT is Shigella toxin and the GL is lactosylceramide, the BT is Shiga toxin and the GL is globoside, digalactosylceramide and/or globotriosyl ceramide, the BT is botulinum toxin and the GL is disialoganglioside or trisialoganglioside or the BT is tetanus toxin and the GL is trisialoganglioside. The pathogen is a bacterium.

USE - The process may be used for **detecting**pathogens including bacteria or viruses, toxins,
lymphocytes, neutrophils, platelets or cancer cells including colon
carcinoma and Burkitt's tumour, in samples, including food, urine,
serum or biopsies.

ADVANTAGE - The PVDF surface is a good substrate for receptor binding assays. The process is highly specific and does not require special equipment. Results can be obtained within a few hours with the naked eye. As the lipid-based receptors have a long shelf-life, they can be easily stored and used for a long period.

Dwg.0/7

L20 ANSWER 12 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

1998-532018 [45] WPIDS

CROSS REFERENCE:

2001-190945 [14]

DOC. NO. CPI:

C1998-159730

TITLE:

Reagent for detecting bacteria and fungi, in e.g.

in food - comprises labelled murein binding

polypeptide and labelled antibiotic for detecting

bound polypeptide.

DERWENT CLASS:

A96 B04 C07 D16

INVENTOR(S):

LAINE, R A; LO, W C J

PATENT ASSIGNEE(S):

(LOUU) UNIV LOUISIANA STATE & AGRIC & MECH COLL; (LAIN-I) LAINE R A; (LOWC-I) LO W C J; (ANOM-N)

ANOMERI INC; (LOUU) UNIV LOUISIANA STATE

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LAPG ______

WO 9842864 A1 19981001 (199845) * EN 110

RW: AT BE CH DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW

NL OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE GH HU IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD

MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR

TT UA UG US UZ VN YU ZW

A 19981020 (199909) AU 9869401 A 19990810 (199938) US 5935804

A1 20000223 (200015) EP 980439

R: CH DE DK FI FR GB IT LI NL SE

A 20000718 (200037) US 6090573 A 20001212 (200067)

US 6159719

JP 2002503093 W 20020129 (200211) 138

APPLICATION DETAILS:

PAT	TENT NO	KIND		API	PLICATION	DATE
WO	9842864	A1			1998-US5580	19980320
ΑŲ	9869401	Α			1998-69401	19980320
US	5935804	Α		US	1997-823293	19970321
ΕP	980439	A1		EP	1998-915148	19980320
				WO	1998-US5580	19980320
US	6090573	Α	Cont of	US	1997-823293	19970321
				US	1999-261664	19990303
US	6159719	Α	Div ex	US	1997-823293	19970321
				US	1999-261665	19990303
JΡ	200250309	3 W		JP	1998-545847	19980320
				WO	1998-US5580	19980320

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9869401 EP 980439	A Based on Al Based on	WO 9842864 WO 9842864
US 6090573	A Cont of	US 5935804

Searcher : Shears

308-4994

A Div ex US 5935804 US 6159719 JP 2002503093 W Based on WO 9842864

PRIORITY APPLN. INFO: US 1997-823293 19970321; US 1999-261664 19990303; US 1999-261665 19990303

1998-532018 [45] WPIDS AN

2001-190945 [14] CR

9842864 A UPAB: 20020215 AΒ

Diagnostic reagent (A) for detecting eubacteria and fungi comprises a murein binding polypeptide (I) conjugated to a reporter (II). (I) is an enzyme having a site that binds eubacterial murein (II) or fungal murein like compounds (III) with binding affinity 5 multiply 10-7-5 multiply 10-9 1/mole, and has substrate turnover rate for (III) or (IIIa) < 3 m mole/minute.

Also new are (1) general method for detecting eubacteria and fungi from binding reaction with (I); (2) kits for this process comprising (A), solution for alkaline hydrolysis and reagent for N-acetylation of sugar residues; (3) diagnostic reagent (A') for detecting (I) bound to a murein in a cell wall consisting of antibiotic (IV) and (II); (4) kits for determining antibiotic sensitivity of many eubacteria and fungi in < 12 hours.

USE - Used to detect and quantify (pathogenic) bacteria and fungi in biological fluids, water, foods, air etc., also to screen for antibiotic resistance.

ADVANTAGE - The reagents can detect small numbers of killed or treated pathogens from a wide range of genera. It does not react with normal mammalian tissue and can differentiate between bacteria and fungi. They have a long shelf live, provide rapid results (usually available within 3 hours), do not require an overnight culture, are suitable for automation and do not need specialised equipment or specifically trained personnel. Dwq.0/12

L20 ANSWER 13 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1992-080211 [10] WPIDS

N1992-060054 DOC. NO. NON-CPI: DOC. NO. CPI: C1992-037175

New adhesion receptors for pathogenic and TITLE:

> opportunistic microorganisms - useful as vaccines and for diagnosis, treatment and prevention of pathogenic and opportunistic infections e.g.

salmonella.

B04 D16 S03 DERWENT CLASS:

KRIVAN, H C; SAMUEL, J E INVENTOR(S):

(ANTE-N) ANTEX BIOLOGICS INC; (BIOC-N) BIOCARB INC; PATENT ASSIGNEE(S):

(MICR-N) MICROCARB INC

COUNTRY COUNT: 17

PATENT INFORMATION:

PG PATENT NO KIND DATE WEEK LA

A 19920220 (199210)* 67 WO 9202817 RW: AT BE CH DE DK ES FR GB GR IT LU NL SE

W: CA JP

A1 19930804 (199331) EN EP 553113 R: AT CH DE DK ES FR GB IT LI NL SE

> Shears 308-4994 Searcher :

JΡ	06501383	W 19	940217	(199412)	19
EΡ	553113	A4 19	940330	(199530)	
ŲS	5696000	A 19	971209	(199804)	20
ΕP	553113	B1 19	981125	(199851)	EN
	R: AT CH	DE DK	ES FR G	GB IT LI NI	SE
DE	69130536	E 19	990107	(199907)	
ES	2127198	T3 19	990416	(199922)	
CA	2095642	C 19	991214	(200018)	EN

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 553113	A1	EP 1991-916508	19910729
JP 06501383	W	WO 1991-US5179 JP 1991-515061	19910729 19910729
		WO 1991-US5179	19910729
EP 553113	A4	EP 1991-916508	
US 5696000	A Div ex	US 1990-562002	19900802
	Cont of	US 1993-78660	19930621
		US 1994-275702	19940718
EP 553113	B1	EP 1991-916508	19910729
		WO 1991-US5179	19910729
DE 69130536	E	DE 1991-630536	19910729
		EP 1991-916508	19910729
		WO 1991-US5179	19910729
ES 2127198	т3	EP 1991-916508	19910729
CA 2095642	C	CA 1991-2095642	19910729
		WO 1991-US5179	19910729

FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 553113	Al Based on	WO 9202817
JP 06501383	W Based on	WO 9202817
EP 553113	B1 Based on	WO 9202817
DE 69130536	E Based on	EP 553113
	Based on	WO 9202817
ES 2127198	T3 Based on	EP 553113
CA 2095642	C Based on	WO 9202817

PRIORITY APPLN. INFO: US 1990-562002 19900802; US 1993-78660 19930621; US 1994-275702 19940718

AN 1992-080211 [10] WPIDS

AB WO 9202817 A UPAB: 19970716

A receptor (I) comprises a substantially pure compound e.g. GalB1-4GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-4GlcNAcB1-3GalB1-4Glc, GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-3GalB1-4Glc, GalB1-3GalB1-4Glc, GalB1-3GalB1-3Gal, where x is sphingosine, hydroxylated sphingosine or saturated sphingosine, and R is H or N-acyl fatty acid derivative of x such that x(R) is a ceramide.

Also new are (1) a compsn. comprising (I) attached to an insoluble or soluble substrate; (2) a method for detecting microorganisms in a sample comprising:- (a) contacting the sample with (I) for a period of time and under conditions sufficient for the receptors to bind the microorganisms; and (b) assaying

for complexes formed; (3) a diagnostic **kit** for the **detection** of **pathogenic** or opportunistic microorganisms comprising the compsn. of (1) and means for detecting or measuring the formation of complexes.

USE/ADVANTAGE - (I) can be used in various forms to detect microorganisms, remove them from liquids and treat infections caused by them. It inhibits their adhesion to mammalian cells if suspended in a liquid which is physiologically compatible with the cells. The adhesion protein has diagnostic and therapeutic applications and is particularly useful as a vaccine. @(67pp Dwg.No.0/4

ABEO EP 553113 A UPAB: 19931118

A receptor (I) comprises a substantially pure compound e.g. GalB1-4GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-3GalB1-4Glc, GalB1-3GalB1-4Glc, GalB1-3GalB1-3Gal, where x is sphingosine, hydroxylated sphingosine or saturated sphingosine, and R is H or N-acyl fatty acid derivative of x such that x(R) is a ceramide.

Also new are (1) a compsn. comprising (I) attached to an insoluble or soluble substrate; (2) a method for detecting microorganisms in a sample comprising:— (a) contacting the sample with (I) for a period of time and under conditions sufficient for the receptors to bind the microorganisms; and (b) assaying for complexes formed; (3) a diagnostic kit for the detection of pathogenic or opportunistic microorganisms comprising the compsn. of (1) and means for detecting or measuring the formation of complexes.

USE/ADVANTAGE - (I) can be used in various forms to detect microorganisms, remove them from liquids and treat infections caused by them. It inhibits their adhesion to mammalian cells if suspended in a liquid which is physiologically compatible with the cells. The adhesion protein has diagnostic and therapeutic applications and is particularly useful as a vaccine.

ABEQ US 5696000 A UPAB: 19980126

A receptor (I) comprises a substantially pure compound e.g. GalB1-4GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-4GlcNAcB1-3GalB1-4Glc, GalB1-4Glc, GalB1-4

Also new are (1) a compsn. comprising (I) attached to an insoluble or soluble substrate; (2) a method for detecting microorganisms in a sample comprising:— (a) contacting the sample with (I) for a period of time and under conditions sufficient for the receptors to bind the microorganisms; and (b) assaying for complexes formed; (3) a diagnostic kit for the detection of pathogenic or opportunistic microorganisms comprising the compsn. of (1) and means for detecting or measuring the formation of complexes.

USE/ADVANTAGE - (I) can be used in various forms to detect microorganisms, remove them from liquids and treat infections caused by them. It inhibits their adhesion to mammalian cells if suspended in a liquid which is physiologically compatible with the cells. The adhesion protein has diagnostic and therapeutic applications and is particularly useful as a vaccine.

Dwg.0/4b

L20 ANSWER 14 OF 14 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1981-50373D [28] WPIDS

TITLE: Reagent for determining esterase antibody

- originating from pathogenic

Streptococcus, includes esterase and protein.

DERWENT CLASS: B04 D16

PATENT ASSIGNEE(S): (DAIN) DAINIPPON PHARM CO LTD

COUNTRY COUNT: 13

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
JP 56058499	Α	19810521	(198128)*		
EP 61546	Α	19821006	(198241)	EN	
R: AT BE	CH	DE FR GB 1	T LI LU N	T SE	
EP 61546	В	19840704	(198427)	EN	
R: DE FR	GB	IT ·			
DE 3164520	G	19840809	(198433)		
US 4592995	Α	19860603	(198625)		
JP 62056980	В	19871128	(198751)		

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
	_		10701010
JP 56058499	A	JP 1979-135400	19791019
EP 61546	A	EP 1981-301377	19810330
US 4592995	A	US 1983-552014	19831116

PRIORITY APPLN. INFO: JP 1979-135400 19791019

AN 1981-50373D [28] WPIDS

AB JP 56058499 A UPAB: 19930915

Reagent comprises (1) esterase (a) origined from pathogenic Streptococcus (2) protein (b) which may bond with antibody (d) to esterase (a) origined from pathogenic Streptococcus and bond with insol. carrier and (3) reagent (c) for determination of esterase (a).

Antiesterase antibody (d) i.e. antibody (d) to esterase (a) origined from pathogenic Streptococcus may be determined in an elapse time such as 1-2 hours to give very quantitatively with a little amt. of sample by means of the reagent according to this invention.

ABEQ EP 61546 B UPAB: 19930915

Reagent kit for determining an antibody against an esterase from pathogenic streptococci comprises (A) an esterase from pathogenic streptococci, (B) protein which can bind to an antibody against the esterase (A) and is bound to an insoluble carrier; and (C) a reagent for measuring an activity of the esterase (A).

(A) may be type A-1 esterase, type A-II esterase, type B esterase or a mixt. The **kit** is useful for serodiagnosis of diseases due to streptococcal infections.

ABEQ US 4592995 A UPAB: 19930915

Reagent **kit** for the determination of an antibody active against esterase from pathogenic streptococci in human blood serum samples comprises (a) a standard soln. of the above esterase; (b) an immobilised protein-A (which binds non-specifically to the

PATENT NO	KIND	APPLICATION	DATE
US 6174733	В1	US 1998-93532	19980608

PRIORITY APPLN. INFO: GB 1997-11941 19970609

AN 2001-146292 [15] WPIDS

CR 2000-181990 [16]; 2000-549271 [49]

AB US 6174733 B UPAB: 20010317

NOVELTY - A detection method for detecting the presence of at least two predetermined materials comprises detecting two predetermined materials, one of which is a control, using a column which has a snare for each of the materials.

DETAILED DESCRIPTION - The method comprises:

- (a) introducing a test sample comprising at least one control material into a column having a snare for each predetermined material, the snare having a capture material which **binds** specifically with the associated predetermined material;
 - (b) washing the test column to remove unbound materials; and
- (c) detecting bound materials on each of the snares, optionally by adding a label for each of the bound materials to form labeled bound materials and detecting labeled bound materials.

INDEPENDENT CLAIM are also included for:

- (1) a column for analyzing at least one pathogen by the method, comprising at least two spatially separated snares, one having a control capture material and the other a pathogen capture material to enable detection of control material and a pathogen of interest; and
 - (2) a kit which comprises:
- (a) a column for analysis of at least one pathogen in which the column has at least two snares. and the snares are separated spatially one from another so that the snares are not in contact with one another, on of the snares having on it a first control capture material for detecting the presence of a first control material, and the other of the snares having on them a pathogen capture material for detecting a pathogen; and
- (b) reagents for detecting the presence of materials selected from:
- (i) reagents for detecting the presence of the control pathogen and the test pathogen; and
- (ii) reagents for detecting the presence of the first control capture material and the pathogen capture material after the first control capture material and the pathogen capture material have been bound and then unbound from the first control material and the pathogen material.
- USE The method is useful to detect pathogens e.g. human pathogens such as tuberculosis, especially by detecting proteins, DNA or RNA in serum. For example, a DNA method may be used to diagnose herpes simplex virus and an RNA method to diagnose HIV. The method may also be useful in veterinary medicine and to detect chemicals such as drugs, carcinogens, pollutants etc.

ADVANTAGE - Unlike previous pathogen detection techniques, the method is rapid and is sensitive to any error in the method because

of the inclusion of a control snare. Dwg.0/11

L13 ANSWER 20 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2001-248306 [26] WPIDS

DOC. NO. NON-CPI: DOC. NO. CPI:

N2001-176898 C2001-075067

TITLE:

Detecting test substances useful for

detecting, e.g. pathogens,

comprises adding test liquid, enzyme-labeled

specific binding substance and

substrate liquid to their respective

addition parts on immunological inspection piece.

DERWENT CLASS:

PATENT ASSIGNEE(S):

B04 D16 S03 (NITL) NITTO DENKO CORP

COUNTRY COUNT:

1

PATENT INFORMATION:

KIND DATE WEEK PATENT NO -----JP 2001013140 A 20010119 (200126)*

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE					
TP 20010131	4 O A	JP 1999-182412	19990628					

PRIORITY APPLN. INFO: JP 1999-182412 19990628

AN 2001-248306 [26] WPIDS

JP2001013140 A UPAB: 20010515 AB

NOVELTY - Detecting test substance (T) comprises using base material having fixed part (I) comprising (T) coupled to first immunological reagent (R1), substrate (S) addition part (AP) (II), enzyme(E)-labeled specific binding substance (III) which couples (R2) and (E), to (T). Test liquid-(III) AP is arranged between (I) and (II). The test liquid, (III), (S)-liquid are added in this order in their respective APs.

DETAILED DESCRIPTION - Detecting (T) involves use of immunological inspection piece which has water absorptive base material on which is present, a fixed part (I) comprising (T) to which a first immunological reagent (R1) is coupled, substrate (S) liquid addition part (II) for adding (S), an enzyme(E)-labeled specific binding substance (III) which couples a second immunological reagent (R2) and enzyme, to (T). The addition part of (III) (1) is arranged between (I) and (II).

An INDEPENDENT CLAIM is also included for a kit for detecting substances.

USE - For detecting test substances such as

ADVANTAGE - The method is rapid and simple. Dwg.0/2

L13 ANSWER 21 OF 47 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.

2002:820 BIOSIS ACCESSION NUMBER: PREV200200000820 DOCUMENT NUMBER:

The type III secretion chaperone LcrH co-operates TITLE:

> 308-4994 Shears Searcher :

with YopD to establish a negative, regulatory loop

for control of Yop synthesis in Yersinia

pseudotuberculosis.

Francis, Matthew S. (1); Lloyd, Scott A.; Wolf-Watz, AUTHOR (S):

(1) Department of Molecular Biology, Umea University, CORPORATE SOURCE:

S-90187, Umea: matthew.francis@cmb.umu.se Sweden

Molecular Microbiology, (November, 2001) Vol. 42, No. SOURCE:

4, pp. 1075-1093. print.

ISSN: 0950-382X.

DOCUMENT TYPE: Article English LANGUAGE:

The enteropathogen Yersinia pseudotuberculosis is a model system AB used to study the molecular mechanisms by which Gram-negative pathogens secrete and subsequently translocate antihost effector proteins into target eukaryotic cells by a common type III secretion systems (TTSS). In this process, YopD (Yersinia outer protein D) is essential to establish regulatory control of Yop synthesis and the ensuing translocation process. YopD function depends upon the non-secreted TTSS chaperone LcrH (low-calcium response H), which is required for presecretory stabilization of YopD. However, as a new role for TTSS chaperones in virulence gene regulation has been proposed recently, we undertook a detailed analysis of LcrH. A lcrH null mutant constitutively produced Yops, even when this strain was engineered to produce wild-type levels of YopD. Furthermore, the YopD-LcrH interaction was necessary to regain the negative regulation of virulence associated genesyops). This finding was used to investigate the biological significance of several LcrH mutants with varied YopD binding potential. Mutated LcrH alleles were introduced in trans into a lcrH null mutant to assess their impact on yop regulation and the subsequent translocation of YopE, a Rho-GTPase activating protein, across the plasma membrane of eukaryotic cells. Two mutants, LcrHK20E, E30G, I31V, M99V, D136G and LcrHE30G lost all regulatory control, even though YopD binding and secretion and the subsequent translocation of YopE was indistinguishable from wild type. Moreover, these regulatory deficient mutants showed a reduced ability to bind YscY in the two-hybrid assay. Collectively, these findings confirm that LcrH plays an active role in yop regulation that might be mediated via an interaction with the Ysc secretion apparatus. This chaperone-substrate interaction presents an innovative means to establish a regulatory hierarchy in Yersinia infections. It also raises the questions as to whether or not LcrH is a true chaperone involved in stabilization and secretion of YopD or a regulatory protein responsible for co-ordinating synthesis of Yersinia virulence determinants. We suggest that LcrH can exhibit both of these activities.

DERWENT INFORMATION LTD WPIDS COPYRIGHT 2002 L13 ANSWER 22 OF 47 2001-024865 [03] WPIDS ACCESSION NUMBER:

C2001-007579 DOC. NO. CPI:

TITLE:

New yfjO polypeptides and polynucleotides encoding the yfj0 polypeptides useful for diagnosing and

staging diseases, and in screening for

antibacterial drugs.

DERWENT CLASS: B04 D16

BISWAS, S; BROWN, J R; BRYANT, A; CHALKER, A F; INVENTOR(S): HOLMES, D J; INGRAHAM, K A; SO, C Y; VAN HORN, S;

> Shears 308-4994 Searcher :

WARREN, R L; ZALACAIN, M; BRYANT, A P

PATENT ASSIGNEE(S): (BISW-I) BISWAS S; (BROW-I) BROWN J R; (BRYA-I)

BRYANT A; (CHAL-I) CHALKER A F; (HOLM-I) HOLMES D J; (INGR-I) INGRAHAM K A; (SOCY-I) SO C Y; (VHOR-I) VAN HORN S; (WARR-I) WARREN R L; (ZALA-I) ZALACAIN

M; (SMIK) SMITHKLINE BEECHAM CORP; (SMIK)

SMITHKLINE BEECHAM PLC

COUNTRY COUNT:

20

PATENT INFORMATION:

PAT	ENT	ИО	F	KINE) DA	ATE	W.	EEK		LA.	P(3					
	200 RW: W:												LU	MC	NL	PT	SE

US 6245542 B1 20010612 (200135)

US 2001023064 A1 20010920 (200156)

APPLICATION DETAILS:

PATENT NO K	IND	APPLICATION	DATE
WO 2000068365 US 6245542 US 2001023064	B1	WO 2000-US11450 US 1999-306276 US 1999-306276 US 2000-735735	20000428 19990506 19990506 20001213

FILING DETAILS:

PATENT NO	KIND	PATENT NO
US 20010230	64 Al Div ex	US 6245542

PRIORITY APPLN. INFO: US 1999-306276 19990506; US 2000-735735 20001213

AN 2001-024865 [03] WPIDS

AB WO 200068365 A UPAB: 20010116

NOVELTY - An isolated polypeptide (I) selected from a polypeptide at least 95 % identical to a 451 residue amino acid sequence (S1) corresponding to yfjO polypeptide (Streptococcus pneumoniae tRNA methyltransferase), a polypeptide having (S1), or a polypeptide encoded by a recombinant polynucleotide (II) having a 1356 base pair sequence (S2), is new. Both sequences fully defined in the specification.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) an isolated polynucleotide selected from:

- (a) a polynucleotide encoding a polypeptide having at least 95 % identity to (I);
- (b) a polynucleotide having at least 95 % identity to (II) or to a polynucleotide encoding (I);
 - (c) a polynucleotide encoding (I);
 - (d) a polynucleotide having (S2);
- (e) a polynucleotide at least 30 nucleotides in length obtained by screening an appropriate library under stringent hybridization conditions with a probe having (S2), or its fragment at least 30 nucleotides in length;
 - (f) a polynucleotide encoding a mature polypeptide expressed by

the yfjO gene comprised in the Streptococcus pneumoniae; or

(g) a complement of any of (a)-(f);

- (2) treating an individual in need of enhanced or expression of or immunological response to (I) by administering an antagonist of (I);
- (3) treating an individual having need to inhibit activity or expression of (I) by administering:

(a) an antagonist of (I);

- (b) a nucleic acid that inhibits the expression of a polynucleotide encoding (I);
- (c) a polypeptide that competes with the polypeptide for its ligand or receptor; or
- (d) a polypeptide that induces an immunological response to (I) in the individual;
- (4) diagnosing or prognosing a disease or a susceptibility to a disease in an individual related to expression or activity of (I) in an individual by:
- (a) determining the presence of mutation in the nucleotide sequence encoding (I) in an organism; or
- (b) analyzing for the presence or amount of the polypeptide expression in a sample from the individual;
- (5) producing (I) by culturing a host cell under conditions sufficient for the production of (I);
- (6) producing a host cell comprising an expression system or its membrane expressing (I) by transforming or transfecting a cell with an expression system comprising a polynucleotide capable of producing (I);
 - (7) a host cell or its membrane expressing (I);

(8) an antibody immunospecific for (I);

- (9) screening compounds that agonize or inhibit the function of (I), comprising:
- (a) measuring the binding of a candidate compound to(I), or fusion proteins of it using a label associated directly or indirectly with the candidate compound;
 - (b) measuring the binding of a candidate compound to

(I) in the presence of a labeled competitor;

- (c) testing if the candidate compound results in a signal generated by activation or inhibition of (I);
- (d) mixing a candidate compound with a solution comprising (I), measuring the activity of (I) in the mixture, and comparing the activity to a standard; or
- (e) detecting the effect of a candidate compound on the production of mRNA encoding (I), and (I) in cells, using e.g. enzyme linked immunosorbent assay (ELISA); and

(10) an agonist or antagonist of (I).

ACTIVITY - Cardiant; neuroprotective; dermatological; ophthalmological; osteopathic; nephrotropic.

No biological data is given.

MECHANISM OF ACTION - yfjO agonists and antagonists.

USE - The polynucleotides and polypeptides are useful as research reagents and materials in the discovery of disease treatments, in the diagnosis, staging and type of infection the pathogen has attained, in determining the response of an infectious organism to drugs, and for screening for antibacterial drugs. These may also be used to assess the binding of small molecule substrates and ligands in cells, cell-free preparations, chemical libraries and natural product mixtures, and to identify agonists and antagonists, which may be used for treating or preventing diseases, such as

infections of the respiratory tract, lower respiratory, cardiac, gastrointestinal, eye, central nervous system, skin, kidney and urinary tract, bone and joint. They may be used to configure screening methods for detecting the effect of added compounds on the production of mRNA and/or polypeptide in cells, and in the prevention of bacterial adhesion to eukaryotic extracellular matrix proteins on in-dwelling devices or to extracellular matrix proteins in wounds. The can be used to block bacterial adhesion between eukaryotic extracellular matrix proteins and bacterial yfj0 proteins that mediate tissue damage, and/or to block the normal progression of pathogenesis in infections initiated other than by the implantation of in-dwelling devices or by other surgical techniques. The polypeptides may further be used identify membrane bound or soluble receptors. The polynucleotides may be used as hybridization probes for RNA, cDNA and genomic DNA to isolate full-length cDNAs and genomic clones encoding yfj0 and to isolate cDNA and genomic clones of other genes that have a high identity to a yfjO gene, in the discovery and development of antibacterial compounds, and to construct antisense sequences to control the expression of the coding sequence of interest. Dwg.0/0

ACCESSION NUMBER:

L13 ANSWER 23 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

2000-664984 [64] WPIDS

DOC. NO. NON-CPI:

N2000-492840

DOC. NO. CPI:

C2000-201437

TITLE:

New 0636 regulator polypeptides and

polynucleotides, useful for screening compounds with antimicrobial activity and diagnosing and

treating pathogen infections.

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

BURNHAM, M K R; LUNSFORD, R D; THROUP, J

PATENT ASSIGNEE(S):

(SMIK) SMITHKLINE BEECHAM CORP

COUNTRY COUNT:

19

PATENT INFORMATION:

KIND DATE WEEK PATENT NO WO 2000059923 A1 20001012 (200064) * EN 38

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: JP

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 20000599	23 A1	WO 2000-US8830	20000330

PRIORITY APPLN. INFO: US 1999-286024 19990405

WPIDS AN 2000-664984 [64]

WO 200059923 A UPAB: 20001209 AB

NOVELTY - An isolated polypeptide comprising a sequence (I) of 233 amino acids, a sequence having at least 95% sequence identity to (I), or is encoded by a recombinant polynucleotide (II) having a sequence of 702 bp, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

> Shears 308-4994 Searcher :

- (1) an isolated polynucleotide or its complements selected from:
- (a) a polynucleotide encoding a polypeptide having at least 95% identity to (I);
- (b) a polynucleotide having at least 95% identity to (II) or to a polynucleotide encoding (I);
- (c) a polynucleotide encoding (I), or a polynucleotide having a specified sequence of 702 bp;
- (d) a polynucleotide of at least 30 nucleotides in length obtained by screening a library with a probe having a specified sequence of 702 bp, or its fragment at least 30 nucleotides in length;
- (e) a polynucleotide encoding a mature polypeptide expressed by the 0636 regulator gene in Staphylococcus aureus;
 - (2) treating an individual:
- (a) in need of enhanced activity or expression of or immunological response to (I) by administering an antagonist to (I);
- (b) having need to inhibit activity or expression of (I) by administering an antagonist of (I), a nucleic acid that inhibits the expression of a polynucleotide encoding (I), a polypeptide that competes with (I) for its ligand, substrate or receptor, or a polypeptide that induces an immunological response to (I) in the individual;
- (3) diagnosing or prognosing (susceptibility to) a disease in an individual related to expression or activity of (I) by:
- (a) determining the presence of a mutation in the nucleotide sequence encoding (I); or
- (b) analyzing for the presence or amount of polypeptide expression in a sample from the individual;
 - (4) producing (I) by culturing a host cell;
- (5) producing a host cell comprising an expression system or its membrane expressing (I) by transforming a cell with a vector containing a polynucleotide capable of producing (I);
 - (6) a host cell or its fragment expressing (I);
 - (7) an antibody immunospecific for (I);
- (8) screening for compounds that agonize or inhibit the function of (I) by:
- (a) measuring the **binding** of a candidate compound to (cells or membranes bearing) the polypeptide or a fusion protein by means of a label associated with the candidate compound, or in the presence of a labeled competitor;
- (b) testing whether the candidate compound results in a signal generated by activation or inhibition of the polypeptide, using detection systems appropriate to the cells or cell membranes bearing the polypeptide;
- (c) mixing a candidate compound with a solution comprising (I) to form a mixture, measuring activity of (I) in the mixture, and comparing the activity of the mixture to a standard; or
- (d) detecting the effect of a candidate compound on the production of (mRNA encoding) (I) in cells, by using e.g. enzyme linked immunoassay; and
 - (9) an agonist or antagonist of (I).
 ACTIVITY Antibiotic; antiulcer.

MECHANISM OF ACTION - None given.

USE - The polynucleotides and polypeptides are useful for screening compounds for antimicrobial activity, which determines their role in pathogenesis of infection, dysfunction and disease, as

research reagents and materials for discovery of treatments of and diagnostics of diseases, for assessing the binding of small molecule substrates and ligands in cells, cell-free preparations, chemical libraries and natural product mixtures, and for identifying membrane-bound or soluble receptors. These may further be used in the prevention of bacterial adhesion to eukaryotic extracellular matrix proteins on in-dwelling devices or in wounds, to block bacterial adhesion between eukaryotic extracellular matrix proteins and bacterial 0636 regulator proteins that mediate tissue damage, and/or to block the normal progression of pathogenesis in infections initiated other than by implantation of in-dwelling devices. The polynucleotides may also be used as hybridization probes for RNA, cDNA and genomic DNA, and in the diagnosis of the stage and type of infection the pathogen has attained, especially infections caused by Staphylococcus aureus. The agonists and antagonists may be used to prevent, ameliorate or correct such infections, dysfunction and disease, which include infections of the upper respiratory tract (e.g. otitis media, bacterial tracheitis, acute epiglottitis, thyroiditis), lower respiratory (e.g. emphysema, lung abscess), cardiac (e.g. infective endocarditis), gastrointestinal (e.g. secretory diarrhea, splenic abscess, stomach ulcer), CNS (e.g. cerebral abscess), eye (e.g. conjunctivitis, keratitis, endophthalmitis), kidney and urinary tract (e.g. epididymitis, intrarenal and perinephric abscess), skin (e.g. impetigo, folliculitis), bone and joint (e.g. septic arthritis, osteomyelitis). Dwg.0/0

ACCESSION NUMBER: 2000-611360 [58] WPIDS

L13 ANSWER 24 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

DOC. NO. NON-CPI:

N2000-452760

DOC. NO. CPI:

C2000-182871

TITLE:

Apparatus for use in shear assay system, useful for determining the adhesion of pathogens under physiological shear stress

conditions.

DERWENT CLASS:

B04 D16 J04 S03

90

INVENTOR(S): PATENT ASSIGNEE(S): BARGATZE, R F; CUTLER, J E; GLEE, P M; PYLE, B

(LIGO-N) LIGOCYTE INC

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000052211 A1 20000908 (200058)* EN 43

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW NL OA PT SD SE SL SZ TZ UG ZW

W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000037132 A 20000921 (200065)

APPLICATION DETAILS:

APPLICATION PATENT NO KIND

WO 2000052211 A1 AU 2000037132 A

WO 2000-US5280 AU 2000-37132

20000301 20000301

FILING DETAILS:

PATENT NO PATENT NO KIND WO 200052211 AU 2000037132 A Based on

PRIORITY APPLN. INFO: US 1999-122215P 19990301

2000-611360 [58] WPIDS ΑN AΒ

WO 200052211 A UPAB: 20001114

NOVELTY - Apparatus comprising an elongated tube with an inner surface adapted to support a substrate, a means for producing a flow of fluid comprising test molecules in the tube, test cellular components or test cells, and a means for monitoring the interaction of the test molecules, test cellular components or test cells with the substrate, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) an apparatus comprising:
- (a) a conduit having an inner surface adapted to support a substrate comprising substrate cells, substrate extracellular matrix proteins, substrate molecules, substrate cellular components or combinations of these;
- (b) a means for producing in the conduit a flow of fluid comprising test molecules, test cellular components or test cells;
- (c) a means for monitoring the interaction of the test molecules, test cellular components or test cells with the substrate cells, substrate extracellular matrix proteins, substrate molecules, substrate cellular components or combinations of these; and
- (2) determining the adhesion of pathogens under physiological shear stress conditions using the above apparatus, comprising:
- (a) introducing test molecules, test cellular components or test cells into the fluid flow; permitting the test molecules, test cellular components or test cells to interact with the substrate cells, substrate extracellular matrix proteins, substrate molecules, substrate cellular components or combinations of these; and
 - (b) monitoring the interactions.
- USE The apparatus can be used in a method for determining the adhesion of pathogens under physiological shear stress conditions. The pathogens can be viruses, bacteria, fungi, protozoa, or parasites (all claimed). The apparatus can be used to study leukocyte interactions with vascular endothelium or purified host ligands under simulated physiological shear.

ADVANTAGE - The apparatus system permits the analysis of complex adhesion behaviors that can occur in simulated physiological shear, unlike prior art methods. The apparatus also eliminates the potential perturbations to microbial binding interactions that occur during the washing steps of prior art assays.

DESCRIPTION OF DRAWING(S) - The diagram shows the basic

components of an in vitro shear system. Dwg.1/6

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DERWENT INFORMATION LTD
L13 ANSWER 25 OF 47 WPIDS COPYRIGHT 2002
                      2000-137031 [12] WPIDS
ACCESSION NUMBER:
                      2000-126669 [09]; 2000-126670 [09]; 2000-126695
CROSS REFERENCE:
                      [09]; 2000-126704 [09]; 2000-126834 [09];
                      2000-137017 [09]; 2000-137019 [09]; 2000-137030
                      [09]; 2000-160622 [09]; 2000-160623 [09];
                      2000-365416 [31]; 2001-015868 [62]; 2001-024697
                      [62]; 2001-040841 [62]; 2001-060891 [62];
                      2001-112369 [08]; 2001-122672 [62]
                      N2000-102434
DOC. NO. NON-CPI:
                      C2000-042070
DOC. NO. CPI:
                      New disposable article comprising a biosensor for
TITLE:
                      detection of specific target biological analytes in
                      bodily waste.
                      B04 D16 D22 F07 J04 P32 P34
DERWENT CLASS:
                      FEDOSOV, Y; KHOMIAKOV, O; KRUCHININ, M; MUSCAT, A;
INVENTOR(S):
                      ROE, D C; FESOSOV, Y
                      (PROC) PROCTER & GAMBLE CO
PATENT ASSIGNEE(S):
                      87
COUNTRY COUNT:
PATENT INFORMATION:
     PATENT NO KIND DATE WEEK LA
                                              PG
     _____
     WO 2000000233 A1 20000106 (200012)* EN 65
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC
            MW NL OA PT SD SE SL SZ UG ZW
         W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES
            FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK
            LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG
            SI SK SL TJ TM TR TT UA UG UZ VN YU ZA ZW
     AU 9948421 A 20000117 (200026)
                   A1 20000503 (200026)
     EP 997125
                                          EN
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK
            NL PT RO SE SI
                  A1 20010418 (200123) EN
     EP 1091773
         R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU NL PT SE
     BR 9911750 A 20010403 (200128)
TW 421587 A 20010211 (200146)
CZ 2000004751 A3 20010815 (200157)
     TW 436281 A 20010528 (200172)
KR 2001053327 A 20010625 (200173)
KR 2001080919 A 20010825 (200215)
APPLICATION DETAILS:
                                    APPLICATION DATE
                KIND
     PATENT NO
                                       WO 1999-US14665 19990629
     WO 2000000233 A1
                                        AU 1999-48421 19990629
     AU 9948421
                 A
                                       EP 1998-120476 19981029
     EP 997125
                   A1
                                       EP 1999-932024
                                                         19990629
     EP 1091773
                  A1
                                       WO 1999-US14665 19990629
                                       BR 1999-11750
                                                          19990629
     BR 9911750
                                        WO 1999-US14665 19990629
                                        TW 1999-110980
                                                          19991201
     TW 421587
                   Α
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CZ 2000004	751 A3	WO 1999-US14665	19990629
		CZ 2000-4751	19990629
TW 436281	A	TW 1999-120445	19991123
KR 2001053	327 A	KR 2000-715070	20001229
KR 2001080	919 A	KR 2001-705320	20010427

FILING DETAILS:

PAT	TENT NO	KIND			PAT	CENT NO
AU	9948421	 A	Based	on	WO	200000233
ΕP	1091773	A1	Based	on	WO	200000233
BR	9911750	Α	Based	on	WO	200000233
CZ	200000475	51 A3	Based	on	WO	200000233

PRIORITY APPLN. INFO: US 1999-299399 19990426; US 1998-90993P 19980629; US 1998-106225 19980629; US 1998-107561 19980629; EP 1998-120476 19981029; US 1999-131073P 19990426

AN 2000-137031 [12] WPIDS CR 2000-126669 [09]; 2000-126670 [09]; 2000-126695 [09]; 2000-126704 [09]; 2000-126834 [09]; 2000-137017 [09]; 2000-137019 [09]; 2000-137030 [09]; 2000-160622 [09]; 2000-160623 [09]; 2000-365416 [31]; 2001-015868 [62]; 2001-024697 [62]; 2001-040841 [62]; 2001-060891 [62]; 2001-112369 [08]; 2001-122672 [62]

AB WO 200000233 A UPAB: 20020306

NOVELTY - A disposable article to be fitted to a wearer, comprising a biosensor including at least one bio-recognition element, is new. The biosensor is adapted to detect a target biological analyte in bodily waste or on the wearer's skin.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for a disposable article to be fitted to a wearer comprising a sensor adapted to detect health markers or nutritional markers in the wearer's bodily waste or on the wearer's skin.

USE - The disposable articles are useful for detecting microorganisms and/or other biomolecules in bodily wastes and/or sensors adapted to detect and/or measure components of feces useful as health and/or nutritional indicators, especially in the form of incontinence briefs, incontinence undergarments, absorbent inserts, nappy holders and liners, disposable bed pads, colostomy bags for a natural or artificial anus, feminine hygienbe garments, tampons, wipes, disposable towels, tissues, bibs, water absorbing articles, oil absorbing articles, spill cleanup bags, desiccant bags, disposable mops, bandages, disposable medical undergarments, disposable plates and cups, disposable food preparation and cutting surfaces, therapeutic wraps, supports and disposable heating pads etc (especially sanitary napkin, a nappy, a training pant, an insert and an adult incontinence device (claimed). Particularly the article is useful for detecting calcium malabsorption which may lead to long term bone mass deficiency.

ADVANTAGE - The biosensors function by providing a means of specifically **binding** and therefore detecting a target biologically active analyte, therefore the biosensor is highly selective.

DESCRIPTION OF DRAWING(S) - The figure is a plan view of a nappy in a flat out stage.
Nappy 20;
Topsheet 24;

Backsheet 26;
Absorbent Core 28;
Side panels 30;
Leg Cuffs 32;
Waist Feature 34; and
Fastening feature 40.
Dwg.1/8

L13 ANSWER 26 OF 47 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 2000:489601 BIOSIS DOCUMENT NUMBER: PREV200000489722

TITLE: Characterization of the basis of lipoprotein (a)

lysine-binding heterogeneity.

AUTHOR(S): Xia, Jiazhi; May, Lorraine F.; Koschinsky, Marlys L.

(1)

CORPORATE SOURCE: (1) Department of Biochemistry, Queen's University,

Kingston, ON, K7L 3N6 Canada

SOURCE: Journal of Lipid Research, (October, 2000) Vol. 41,

No. 10, pp. 1578-1584. print.

ISSN: 0022-2275.

DOCUMENT TYPE: Article
LANGUAGE: English
SUMMARY LANGUAGE: English

Although elevated plasma concentrations of lipoprotein (a) (Lp(a)) are considered to be a risk factor for atherosclerosis, the mechanisms by which Lp(a) mediates its pathogenic effects have not been conclusively determined. The apolipoprotein (a) (apo(a)) component of Lp(a) confers unique structural properties to this lipoprotein, including the ability to bind to lysine residues in biological substrates. It has been shown, however, that only a fraction of plasma Lp(a) (Lp(a)-Lys+) binds to lysine-Sepharose in vitro. The nature of the non-lysine-binding Lp(a) fraction in plasma (Lp(a)-Lys-) is currently unknown. In the present study, the Lp(a)-Lys+ fraction was determined in the plasma of six unrelated individuals; the Lp(a)-Lys+ fraction in these plasma samples ranged from apprx37 to apprx48%. Interestingly, purification of the Lp(a) by density gradient ultracentrifugation followed by gel filtration and ion-exchange chromatography resulted in progressive increases in the Lp(a)-Lys+ fraction. Addition of either purified low density lipoprotein (LDL) or fibronectin to the purified Lp(a) at a 1:1 molar ratio reduced the Lp(a)-Lys+ fraction (maximal decrease of 34 and 20%, respectively) whereas addition of both fibronectin and LDL to the purified Lp(a) resulted in a further decrease (45% maximally) in this fraction. Similar results were obtained by using a recombinant expression system for apo(a): addition of a 4-fold molar excess of either LDL or fibronectin to conditioned medium containing metabolically labeled recombinant apo(a) reduced the Lys+ fraction by 49 and 23%, respectively. Taken together, our data suggest that the lysine-binding heterogeneity of plasma Lp(a) is not primarily an intrinsic property of the lipoprotein, but rather results in large part from its ability to noncovalently associate with abundant plasma components such as LDL and fibronectin. These interactions appear to mask the lysine-binding site in apo(a) kringle IV type 10, which mediates the interaction of Lp(a) with lysine-Sepharose. The contribution of these interactions to the function of Lp(a) in vivo remains to be investigated.

L13 ANSWER 27 OF 47 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 2000:229045 BIOSIS DOCUMENT NUMBER: PREV200000229045

TITLE: Comparative analysis of the binding of

antibodies prepared against the insect Spodoptera exigua and against the mycopathogen Nomuraea rileyi. Pendland, Jacquelyn C. (1); Boucias, Drion G. (1)

CORPORATE SOURCE: (1) Entomology and Nematology Department, University

of Florida, Gainesville, FL, 32611-0620 USA

SOURCE: Journal of Invertebrate Pathology, (Feb., 2000) Vol. 75, No. 2, pp. 107-116.

ISSN: 0022-2011.

DOCUMENT TYPE: Article
LANGUAGE: English
SUMMARY LANGUAGE: English

AUTHOR(S):

AB Polyclonal antibodies were produced in mice against Spodoptera exigua (beet armyworm) larval hemolymph and hemocytes and against cell wall surfaces of hyphal bodies and hyphae of the entomopathogenic hyphomycete Nomuraea rileyi. In addition to exhibiting strong activity against their original antigenic substrates, all of the antibodies cross-react extensively

with other substrates. The hemolymph antibody binds to hemocytes and vice versa, and both antibodies cross-react to the insect fat body basement membrane (extracellular matrix (ECM) and to N. rileyi and Beauveria bassiana (another entomopathogenic fungus) cell wall surfaces (ECM). Likewise, the anti-fungal antibodies cross-react with S. exigua hemolymph and hemocytes, especially the granules that may contain ECM components, and with fat body basement membrane. These cross-reactivities are specific as indicated by negative controls in the microscopy and Western blotting assays. Parallel labeling experiments using Con A suggest that the reactive epitopes contain mannose; however, none of the antibodies bind to mannose residues of

nonentomopathogenic Candida albicans or Saccharomyces cerevisiae yeast cells. Thus, these cross-reactivities suggest that the host mimicry expressed by surface components of entomopathogenic fungi represents an important pathogenic determinant.

L13 ANSWER 28 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-147216 [13] WPIDS

CROSS REFERENCE: 2000-160573 [08] DOC. NO. CPI: C2000-046073

TITLE: Assay for determining relative viability

of mutant pathogens in presence of

therapeutics, for designing treatments for human

immune deficiency virus infections.

DERWENT CLASS: B04 D16

INVENTOR(S): ERICKSON, J W; GULNIK, S V

PATENT ASSIGNEE(S): (USSH) US DEPT HEALTH & HUMAN SERVICES; (USSH) US

DEPT HEALTH & SOCIAL SERVICES

COUNTRY COUNT: 85

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 9967417 A2 19991229 (200013)* EN 118

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC
MW NL OA PT SD SE SL SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZW

A 20000110 (200025) AU 9948280

A2 20010404 (200120) EN EP 1088098

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9967417 AU 9948280 EP 1088098	A2 A A2	WO 1999-US14119 AU 1999-48280 EP 1999-931861 WO 1999-US14119	19990623 19990623 19990623 19990623

FILING DETAILS:

PATENT NO		PATENT NO
		
AU 9948280	A Based on	WO 9967417
EP 1088098	A2 Based on	WO 9967417

PRIORITY APPLN. INFO: US 1998-90393P 19980623

2000-147216 [13] WPIDS

CR 2000-160573 [08]

9967417 A UPAB: 20010410 AB

NOVELTY - Assay for determining the biochemical fitness of a biochemical target (BT) in a mutant replicating entity, relative to its predecessor, comprises comparing the biochemical viability (BV) of BT from the mutant and predecessor, in presence of a compound that inhibits BT of the predecessor.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(a) administering a drug (A) that inhibits BT of a pathogen by:

- (i) determining first BV for the pathogen and a mutant that has evolved from it, in presence of first compound (C1) able to inhibit BT;
 - (ii) determining second BV for pathogen and

mutant in presence of at least one additional compound (C2);

- (iii) determining, and comparing, biochemical fitness from (i) and (ii), and
- (iv) administering which of C1 and C2 provides the lower value for biochemical fitness, so that the pathogen is less likely to develop resistance in presence of the selected compound;
- (b) continuous fluorogenic assay for measuring anti-HIV (human immune deficiency virus) protease activity of a protease inhibitor, and
- (c) preventing development of drug resistance in an HIV-infected mammal by administering a drug-resistance inhibitor of formula (II), or its salt, prodrug or ester:

A = bicyclic system;

X = bond, CHR10, CHR10CH2, CH2CHR10, oxygen, NR10 or sulfur; R10 = hydrogen , alkyl, alkenyl or alkynyl;

Q and W = carbonyl, thiocarbonyl or sulfonyl; R2 = hydrogen, 1-6C alkyl, or 2-6 C alkenyl or alkynyl; m = 0-6;

R3 = optionally substituted cycloalkyl, heterocycloalkyl, aryl or heteroaryl;

R4 = hydroxy, oxo, amino or methylamino;

R5 = as R2 or (CH2)qR14;

R14 = cycloalkyl, heterocycloalkyl, aryl or heteroaryl, optionally substituted by at least one halo, hydroxy, methoxy, amino, nitro, thiol or cyano;

R6 = optionally substituted cycloalkyl, heterocycloalkyl, aryl or heteroaryl; or R5, R6 and N-W together comprise a 12-18 membered ring containing at least one additional heteroatom such that a mutant virus, able to develop in the mammal, has lower fitness, relative to its predecessor, in presence of (II).

ACTIVITY - Antiviral; antibacterial; antimalarial; anticancer; anthelmintic.

MECHANISM OF ACTION - Enzyme or other protein inhibition.

USE - The method is used to identify compounds that are least likely to cause development of resistance when used to treat viral (specifically HIV), malarial or bacterial infections or cancer.

ADVANTAGE - The method allows selection of drugs that have the best chance of providing successful long-term therapy. The new fluorogenic assay, used to determine viability of HIV-1 in presence of protease inhibitors, is more sensitive than known methods, and is especially useful with multidrug-resistant mutants. The compounds (II) for preventing development of resistant mutants has a strong, broad-spectrum of inhibition against a panel of mutant HIV proteases. Dwq.0/5

L13 ANSWER 29 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-116792 [10] WPIDS

N2000-088408 DOC. NO. NON-CPI:

C2000-035742 DOC. NO. CPI:

Rapid production of chemiluminescence comprises TITLE: e.g. contacting an acid with an acridan compound and contacting the product formed with base to give

a second product with formation of light.

B02 B04 D13 D16 S03 DERWENT CLASS:

AKHAVAN-TAFTI, H; TAFTI-AKHAVAN, H INVENTOR(S):

PATENT ASSIGNEE(S): (LUMI-N) LUMIGEN INC

23 COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 9966328 A1 19991223 (200010) * EN 79

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: AU CA JP

US 6017769 A 20000125 (200012) AU 9937408 A 20000105 (200024) EP 1005649 A1 20000607 (200032)

EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

APPLICATION DATE PATENT NO KIND

WO	9966328	A1	WO	1999-US6560	19990503
US	6017769	A	US	1998-99656	19980617
ΑU	9937408	A	ΑU	1999-37408	19990503
EΡ	1005649	A1	EP	1999-919758	19990503
			WO	1999-US6560	19990503

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9937408	A Based on	WO 9966328
EP 1005649	Al Based on	WO 9966328

PRIORITY APPLN. INFO: US 1998-99656 19980617

AN 2000-116792 [10] WPIDS

AB WO 9966328 A UPAB: 20000228

NOVELTY - Producing chemiluminescence comprises (a) contacting an acid with an acridan compound to form a 1st reaction product (1); and (b) contacting (1) with sufficient quantity of base to provide a basic environment so that a 2nd reaction product (2) is formed and light is produced in the basic environment. At least one of (a) and (b) provides an oxidant for the reaction with (I) or (1).

DETAILED DESCRIPTION - Producing chemiluminescence comprises:

- (a) contacting an acid with an acridan of formula (I) to form a 1st reaction product (1); and
- (b) contacting (1) with sufficient quantity of base to provide A basic environment so that a 2nd reaction product (2) is formed and light is produced in the basic environment. At least one of (a) and (b) provides an oxidant for the reaction with (I) or (1).

Z1, Z2 = 0, S or NR12;

R12 = alkyl, aryl, alkylsulfonyl or arylsulfonyl;

- R1 = 1-50 non-H atoms chosen from C, N, O, S, P, Si or halo that are removable by acid; and
- R2, R3 = organic groups containing 1-50 non-H atoms chosen from C, N, O, S, P, Si or halo; and

R4-R11 = H or substituents that do not interfere with generation of chemiluminescence.

An INDEPENDENT CLAIM is also included for a **method** of performing an assay of analyte in a sample comprising:

(i) generating chemiluminescence from (I);

(ii) detecting the chemiluminescence;

(iii) relating the chemiluminescence to the amount of the

analyte in the sample.

USE - To produce chemiluminescence (claimed). To perform assay of analyte in samples such as drugs, hormones, pesticides, metabolites, DNA, RNA, oligonucleotides, antibodies, antibody fragments, antibody-DNA chimeras, antigens, haptens, proteins, carbohydrates, lectins and receptors (claimed). To detect analytes, especially within electrophoresis gels and in immunoassays, nucleic acid probes, nucleic acid hybridization probes, other ligand-binder assays, western blot, northern blot, southern blot, DNA sequence analysis, colony hybridization, gene expression analysis, high-throughput drug screening, and detection of infectious agents and pathogens, and for detection of analytes in food, environmental and industrial samples. Used for signaling, emergency lighting and novelty items.

Reagent comprising 3.3 multiply 10-4M acridan phosphate in 0.1M tris-(hydroxymethyl)aminomethane (Tris) buffer, pH 8.8 (10 micro 1)

was mixed with 3.6% urea peroxide (50 micro 1) in 0.4M nitric acid and was incubated for 2 minutes. Chemiluminescence was triggered by injecting 0.25M sodium hydroxide solution (100 micro 1). Light production occurred instantly upon mixing and was integrated for 5 seconds.

ADVANTAGE - Rapidly produces chemiluminescence from electron-rich alkenes by simple chemical process using inexpensive, readily available reagents. Detects very small amounts of compounds due to either low abundance in the sample or limited sample quantity. Detects quantity of compound precisely over wide range of concentrations. Dwg.0/3

L13 ANSWER 30 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

2000-147108 [13] WPIDS ACCESSION NUMBER:

N2000-108903 DOC. NO. NON-CPI: C2000-045999 DOC. NO. CPI:

Rapid and automatic system for detection of ligands TITLE:

using highly specific receptors.

B04 D16 J04 S03 DERWENT CLASS:

DOANE, K J; LAVRENTOVICH, O; NIEHAUS, G D; SCHMIDT, INVENTOR(S):

S P; SIGNS, S A; WOOLVERTON, C J

(UYKE-N) UNIV KENT STATE PATENT ASSIGNEE(S):

COUNTRY COUNT: 87

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
WO 9964862			(200013)*		41

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW NL OA PT SD SE SL SZ UG ZW

W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG UZ VN YU ZA ZW

AU 9939844 A 19991230 (200022)

US 6171802 B1 20010109 (200104) BR 9910982

A 20010213 (200114) A1 20010328 (200118) EN EP 1086374

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9964862 AU 9939844	A1 A	WO 1999-US10413 AU 1999-39844	19990512 19990512
US 6171802 BR 9910982	B1 A	US 1998-95196 BR 1999-10982	19980610 19990512
EP 1086374	A1	WO 1999-US10413 EP 1999-922970 WO 1999-US10413	19990512 19990512 19990512
		MO 1333-0210413	19990012

FILING DETAILS:

PATENT NO	KIND	FATENT NO
AU 9939844	A Based on	WO 9964862

BR 9910982 A Based on WO 9964862 EP 1086374 Al Based on WO 9964862

PRIORITY APPLN. INFO: US 1998-95196 19980610

AN 2000-147108 [13] WPIDS

AB WO 9964862 A UPAB: 20000313

NOVELTY - A system for the detection of ligands comprises at least one receptor and an amplification mechanism coupled to the receptor. The amplification mechanism generates a signal upon receptor-ligand binding.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for a device for the detection and monitoring of the presence of ligands comprising multiple wells, each well containing a predetermined receptor coupled to an amplification mechanism. Upon binding of a specific ligand the predetermined receptor is activated and generates a signal, which is then amplified.

USE - The system/device is useful for the detection of ligand by a receptor, especially pathogens and/or toxins.

ADVANTAGE - The method uses highly specific receptors for the rapid and automatic detection of the ligand therefore providing a system for the early detection of pathogenic agents. The system is compact enough to be placed in a physicians office to enable immediate diagnoses.

Dwg.0/7

L13 ANSWER 31 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2000-053261 [04] WPIDS

DOC. NO. CPI:

C2000-013910

TITLE:

Detecting a target nucleic acid fragment in a

clinical specimen, nucleic acid probes for Babesia

species and Borrelia burgdorferi.

DERWENT CLASS:

B04 C07 D16

INVENTOR(S):

HARRIS, N S; SHAH, J S

PATENT ASSIGNEE(S):

(IGEN-N) IGENEX INC

COUNTRY COUNT:

21

PATENT INFORMATION:

PATENT	NO	KIND	DATE	WEEK	LA	PG

WO 9960009 A1 19991125 (200004)* EN 49 RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: AU CA JP

AU 9940011 A 19991206 (200019)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
	A1 A	WO 1999-US10939 AU 1999-40011	

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AII 9940011	A Based	on WO 9960009

PRIORITY APPLN. INFO: US 1998-88696P 19980521; US 1998-88541P 19980521

AN 2000-053261 [04] WPIDS

AB WO 9960009 A UPAB: 20000124

NOVELTY - A method for detecting a target nucleic acid fragment in a clinical specimen obtained from a patient comprises a two part method comprising isolating one or more nucleic acid fragments of the specimen via hybridization with a probe complex, and detection of the isolated nucleic acid fragment.

DETAILED DESCRIPTION - A method for detecting a target nucleic acid fragment in a clinical specimen obtained from a patient comprises:

- (a) solubilizing a sample of the clinical specimen in a chaeotropic salt solution;
- (b) treating the solubilized sample by means to denature the nucleic acids contained in the sample;
- (c) contacting the solubilized sample of (b) with at least one probe complex, the probe complex comprising a nucleic acid sequence which is complementary to a portion of the target nucleic acid fragment and further comprising a first member of a specific binding pair;
- (d) incubating the solubilized sample with probe complex under conditions appropriate for hybridization of the probe complex with the target nucleic acid fragment;
- (e) contacting the probe complex in the incubated sample of (d) with a solid substrate which is linked to the second member of the specific binding pair under conditions which promote binding of the specific binding pair, to isolate hybridized target nucleic acid fragment in probe-target-solid substrate ternary complex;
- (f) separating the isolated probe-target-solid substrate complex from the solubilized sample;
- (g) releasing the target nucleic acid and the probe complex into solution from the separated probe-target-solid substrate complex; amplifying the released target nucleic acid fragment by PCR or RT-PCR; and
- (h) detecting the presence of the target nucleic acid fragment in the clinical specimen by comparison of the amplification products produced by (g) to amplification products produced in identically treated positive and negative control reactions.

INDEPENDENT CLAIMS are also included for the following:

- (1) a method for direct amplification of a target nucleic acid fragment by PCR in a reaction buffer of 30 mM Tris-HCl (pH 8.4), 50 mM KCl, 1. 5 mM MgCl2, by otherwise standard procedures;
- (2) a nucleic acid probe or primer for the detection of Babesia microti or B. WA-1, the probe consisting essentially of at least about 10 consecutive nucleotides of the nucleotide sequence of B5, B6, B7, B8-1 or their complements: and a method for specifically amplifying B. microti nucleic acid sequences by PCR or RT-PCR; and
- (3) a method for detecting Babesia specific nucleic acids in a sample by hybridizing to one of the oligonucleotides in (2) or B8-2:
- USE The methods provide a means of detecting a target nucleic acid fragment in a clinical sample. In particular, the methods are useful for detecting pathogens such as Borrelia burgdorferi and species of

Babesia, the causative agents of Lyme disease and babesiosis (which is common in many wild and domestic animals, less so in humans).

ADVANTAGE - Isolation of target nucleic acid fragments removes unwanted nucleic acids from the desired target nucleic acid fragment, and also significantly eliminates PCR inhibitors from the target nucleic acid fragment. This allows for the highly sensitive and accurate detection of the isolated target nucleic acid fragment. The methods can purify and concentrate several different DNA fragments from a sample in the same reaction, decreasing the total size of a sample required for the performance of multiple analyses. Dwq.0/0

L13 ANSWER 32 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

1999-179969 [15] WPIDS ACCESSION NUMBER:

1996-383666 [38] CROSS REFERENCE: C1999-052344

DOC. NO. CPI: Concurrent processing of multiple biological chip TITLE:

assays - using biological chip plate bearing

multiple probe arrays.

A96 B04 D16 DERWENT CLASS:

FODOR, S P; RAVA, R P; TRULSON, M INVENTOR(S):

(AFFY-N) AFFYMETRIX INC PATENT ASSIGNEE(S):

COUNTRY COUNT:

PATENT INFORMATION:

PATEN'	T NO	KIND	DATE	WEEK .	LA	PG
						
US 58	74219	Α	19990223	(199915)*	•	17

APPLICATION DETAILS:

PATENT NO KIND		APPLICATION	DATE
US 5874219	_	US 1995-476850 US 1996-630051	19950607

FILING DETAILS:

PATENT NO	KIND	PATENT NO
US 5874219	A Cont of	US 5545531

PRIORITY APPLN. INFO: US 1995-476850 19950607; US 1996-630051 19960409

1999-179969 [15] WPIDS AN

CR 1996-383666 [38]

5874219 A UPAB: 19990416 AB HS The following are claimed:

(1) a method for concurrently processing multiple

biological chip assays, comprising:

- (a) providing a biological chip plate comprising a contiguous substrate that comprises a contiguous surface and, attached to the contiguous surface, several probe arrays and, surrounding the probe arrays, material resistant to the flow of liquid, thereby forming several test wells, each test well defining a space for the introduction of a sample;
 - (b) manipulating the biological chip plate with a fluid

308-4994 Searcher : Shears

handling device that automatically performs steps to carry out reactions between target molecules in a test sample and probes in several of the test wells, and

- (c) interrogating the probe arrays of the biological chip plate with a biological chip plate reader to detect reactions between target molecules and probes in several of the test wells to generate assay results;
- (2) a system for concurrently processing multiple biological chip assays, comprising:
- (a) a biological chip plate comprising a contiguous substrate that comprises a contiguous surface and, attached to the contiguous surface, several probe arrays and, surrounding the probe arrays, material resistant to the flow of liquid, thereby forming several test wells, each test well defining a space for the introduction of a sample;
- (b) a fluid handling **device** that automatically performs steps to carry out reactions between target molecules in the samples and probes in several of the test wells; and
- (c) a biological chip plate reader that interrogates the probe arrays to detect any reactions between target molecules and probes in several of the test wells to produce assay results;
- (3) a biological chip plate comprising a contiguous substrate that comprises on its surface several probe arrays and, surrounding the probe arrays, material resistant to the flow of liquid thereby forming several test wells, each test well defining a space for the introduction of a sample;
- (4) a method for making a biological chip plate, comprising providing a contiguous substrate and a body, the contiguous substrate comprising a contiguous surface and, attached to the contiguous surface, several probe arrays, and the body comprising several channels; and attaching the body to the surface of the contiguous substrate, whereby several of the channels each cover a probe array and the surface of the substrate closes one end of the channels, thereby forming test wells defining spaces for receiving samples, and
- (5) a wafer comprising a contiguous substrate that comprises a contiguous surface and, attached to the contiguous surface, several probe arrays, where the probe arrays are arranged on the wafer surface in rows and columns, where the probe arrays in each row are the same and the probe arrays in each column are different.

USE - The assay is useful for:

(a) the detection or identification of a

pathogenic organism, especially HIV;

(b) the detection or identification of a human nucleic acid, preferably a human gene variant, especially where the human gene variant indicates the existence of, or predisposition to cystic fibrosis, diabetes, muscular dystrophy or cancer, and

(c) the identification of a probe in a library that ${\bf binds}$ to a receptor. ${\rm Dwg.0/8}$

L13 ANSWER 33 OF 47 MEDLINE

ACCESSION NUMBER: 1999443902 MEDLINE

DOCUMENT NUMBER: 99443902 PubMed ID: 10512717

TITLE: Crystal structure of an Fab fragment in

complex with a meningococcal serosubtype antigen and a protein G domain.

Derrick J P; Maiden M C; Feavers I M AUTHOR:

Department of Biomolecular Sciences, UMIST, CORPORATE SOURCE:

Manchester, M60 1QD, UK.. Jeremy.Derrick@umist.ac.uk JOURNAL OF MOLECULAR BIOLOGY, (1999 Oct 15) 293 (1)

SOURCE: 81-91.

Journal code: J6V; 2985088R. ISSN: 0022-2836.

ENGLAND: United Kingdom PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

Priority Journals FILE SEGMENT:

OTHER SOURCE: PDB-1QKZ ENTRY MONTH: 199911

Entered STN: 20000111 ENTRY DATE:

Last Updated on STN: 20000111 Entered Medline: 19991122

Many pathogens present highly variable surface proteins to their AΒ host as a means of evading immune responses. The structure of a peptide antigen corresponding to the subtype P1.7 variant of the porin PorA from the human pathogen Neisseria meningitidis was determined by solution of the X-ray crystal structure of the ternary complex of the peptide (ANGGASGQVK) in complex with a Fab fragment and a domain from streptococcal protein

G to 1.95 A resolution. The peptide adopted a beta-hairpin structure with a type I beta-turn between residues Gly4P and Gly7P, the conformation of the peptide being further stabilised by a pair of hydrogen bonds from the side-chain of Asn2P to main-chain atoms in Val9P. The antigen binding site within the Fab formed a distinct crevice lined by a high proportion of apolar amino acids. Recognition was supplemented by hydrogen bonds from heavy chain residues Thr50H, Asp95H, Leu97H and Tyr100H to main-chain and side-chain atoms in the peptide. Complementarity-determining region (CDR) 3 of the heavy chain was responsible for approximately 50 % of the buried surface area formed by peptide-Fab binding, with the remainder made up from CDRs 1 and 3 of the light chain and CDRs 1 and 2 of the heavy chain. Knowledge of the structures of variable surface antigens such as PorA is an essential prerequisite to a molecular understanding of antigenic variation and its implications

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L13 ANSWER 34 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1998-532018 [45]

for vaccine design.

2001-190945 [14] CROSS REFERENCE:

C1998-159730 DOC. NO. CPI:

Reagent for detecting bacteria and fungi, in e.g. TITLE: in food - comprises labelled murein binding

polypeptide and labelled antibiotic for detecting

WPIDS

bound polypeptide. A96 B04 C07 D16

DERWENT CLASS: INVENTOR(S):

LAINE, R A; LO, W C J (LOUU) UNIV LOUISIANA STATE & AGRIC & MECH COLL; PATENT ASSIGNEE(S):

(LAIN-I) LAINE R A; (LOWC-I) LO W C J; (ANOM-N) ANOMERI INC; (LOUU) UNIV LOUISIANA STATE

80 COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK

A1 19981001 (199845)* EN 110 WO 9842864 RW: AT BE CH DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE GH HU IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZW A 19981020 (199909) AU 9869401 A 19990810 (199938) US 5935804 EP 980439 A1 20000223 (200015) EN R: CH DE DK FI FR GB IT LI NL SE A 20000718 (200037) US 6090573 A 20001212 (200067) US 6159719 138 JP 2002503093 W 20020129 (200211)

APPLICATION DETAILS:

PATENT NO F	KIND	APPLICATION	DATE
WO 9842864 AU 9869401 US 5935804 EP 980439	A1 A A A1	WO 1998-US5580 AU 1998-69401 US 1997-823293 EP 1998-915148	19980320 19980320 19970321 19980320 19980320
US 6090573	A Cont of	WO 1998-US5580 US 1997-823293 US 1999-261664	19970321 19990303
US 6159719	A Div ex	US 1997-823293 US 1999-261665	19970321 19990303
JP 2002503093	3 W	JP 1998-545847 WO 1998-US5580	19980320 19980320

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9869401 EP 980439 US 6090573 US 6159719	A Based on Al Based on A Cont of A Div ex	WO 9842864 WO 9842864 US 5935804 US 5935804
*	3 W Based on	WO 9842864

PRIORITY APPLN. INFO: US 1997-823293 19970321; US 1999-261664 19990303; US 1999-261665 19990303

AN 1998-532018 [45] WPIDS

CR 2001-190945 [14]

AB WO 9842864 A UPAB: 20020215

Diagnostic reagent (A) for detecting eubacteria and fungi comprises a murein binding polypeptide (I) conjugated to a reporter (II). (I) is an enzyme having a site that binds eubacterial murein (II) or fungal murein like compounds (III) with binding affinity 5 multiply 10-7-5 multiply 10-9 l/mole, and has substrate turnover rate for (III) or (IIIa) < 3 m mole/minute.

Also new are (1) general method for detecting eubacteria and fungi from binding reaction with (I); (2) kits for this process comprising (A), solution for alkaline hydrolysis and reagent for N-acetylation of sugar residues; (3) diagnostic reagent (A') for detecting (I) bound to a murein in a

cell wall consisting of antibiotic (IV) and (II); (4) kits for determining antibiotic sensitivity of many eubacteria and fungi in < 12 hours.

USE - Used to detect and quantify (pathogenic) bacteria and fungi in biological fluids, water, foods, air etc., also to screen for antibiotic resistance.

ADVANTAGE - The reagents can detect small numbers of killed or treated pathogens from a wide range of genera. It does not react with normal mammalian tissue and can differentiate between bacteria and fungi. They have a long shelf live, provide rapid results (usually available within 3 hours), do not require an overnight culture, are suitable for automation and do not need specialised equipment or specifically trained personnel. Dwq.0/12

BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L13 ANSWER 35 OF 47

ACCESSION NUMBER: DOCUMENT NUMBER:

CORPORATE SOURCE:

1998:428125 BIOSIS PREV199800428125

TITLE:

Cloning and characterization of mitochondrial

methionyl-tRNA synthetase from a pathogenic fungi

Candida albicans.

AUTHOR(S):

Lee, Sang Won; Jo, Yeong Joon; Kim, Sunghoon (1)

(1) Dep. Biol., Sung Kyun Kwan Univ., 300

Chunchundong, Jangangu, Suwon, Kyunggido 440-746

South Korea

SOURCE:

Gene (Amsterdam), (July 30, 1998) Vol. 215, No. 2,

pp. 311-318.

ISSN: 0378-1119.

DOCUMENT TYPE:

Article English

LANGUAGE: A genomic sequence encoding mitochondrial methionyl-tRNA synthetase AB (MetRS) was determined from a pathogenic fungi Candida albicans. The gene is distinct from that encoding the cytoplasmic MetRS. The encoded protein consists of 577 amino acids (aa) and contains the class I defining sequences in the N-terminal domain and the conserved anticodon-binding amino acid, Trp, in the C-terminal domain. This protein showed the highest similarity with the mitochondrial MetRSs of Saccharomyces cerevisiae and Schizosaccharomyces pombe. The mitochondrial MetRSs of these fungi were distinguished from their cytoplasmic forms. The protein lacks the zinc binding motif in the N-terminal domain and the C-terminal dimerization appendix that are present in MetRSs of several other species. Escherichia coli tRNAMet was a substrate for the encoded protein as determined by genetic complementation and in vitro aminoacylation reaction. This cross-species aminoacylation activity suggests the conservation of

DUPLICATE 1 L13 ANSWER 36 OF 47 MEDLINE

ACCESSION NUMBER:

1999030465 MEDLINE

interaction mode between tRNAMet and MetRS.

DOCUMENT NUMBER:

PubMed ID: 9811546 99030465 The crystal structure of the L1

TITLE:

metallo-beta-lactamase from Stenotrophomonas

maltophilia at 1.7 A resolution.

AUTHOR:

Ullah J H; Walsh T R; Taylor I A; Emery D C; Verma C

S; Gamblin S J; Spencer J

CORPORATE SOURCE:

Division of Protein Structure, National Institute of Medical Research, The Ridgeway, Mill Hill, London,

308-4994 Searcher : Shears

NW7 1AA, UK.

SOURCE: JOURNAL OF MOLECULAR BIOLOGY, (1998 Nov 20) 284 (1)

125-36.

Journal code: J6V; 2985088R. ISSN: 0022-2836.

PUB. COUNTRY: ENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT:

OTHER SOURCE:

Priority Journals
PDB-1SML; PDB-R1SMLSF

ENTRY MONTH: 199902

ENTRY DATE: Entered STN: 19990311

Last Updated on STN: 19990311 Entered Medline: 19990225

The structure of the L1 metallo-beta-lactamase from the AB opportunistic pathogen Stenotrophomonas maltophilia has been determined at 1.7 A resolution by the multiwavelength anomalous dispersion (MAD) approach exploiting both the intrinsic binuclear zinc centre and incorporated selenomethionine residues. L1 is unique amongst all known beta-lactamases in that it exists as a tetramer. The protein exhibits the alphabeta/betaalpha fold found only in the metallo-beta-lactamases and displays several unique features not previously observed in these enzymes. These include a disulphide bridge and two substantially elongated loops connected to the active site of the enzyme. Two closely spaced zinc ions are bound at the active site with tetrahedral (Zn1) and trigonal bipyramidal (Zn2) co-ordination, respectively; these are bridged by a water molecule which we propose acts as the nucleophile in the hydrolytic reaction. Ligation of the second zinc ion involves both residues and geometry which have not been previously observed in the metallo-beta-lactamases. Simulated binding of the substrates ampicillin, ceftazidime and imipenem suggests that the substrate is able to bind to the enzyme in a variety of different conformations whose common features are direct interactions of the beta-lactam carbonyl oxygen and nitrogen with the zinc ions and of the beta-lactam carboxylate with Ser187. We describe a catalytic mechanism whose principal features are a nucleophilic attack of the bridging water on the beta-lactam carbonyl carbon, electrostatic stabilisation of a negatively charged tetrahedral transition state and protonation of the beta-lactam nitrogen by a second water molecule co-ordinated by Zn2. Further, we propose that direct metal:substrate interactions provide a substantial contribution to substrate binding and that this may explain the lack of specificity which is a feature of this class of enzyme. Copyright 1998 Academic Press

L13 ANSWER 37 OF 47 SCISEARCH COPYRIGHT 2002 ISI (R)

ACCESSION NUMBER: 97:792093 SCISEARCH

THE GENUINE ARTICLE: YC180

TITLE: Crystal structure of the

phosphatidylinositol-specific phospholipase C from

the human pathogen Listeria monocytogenes

AUTHOR: Moser J; Gerstel B; Meyer J E W; Chakraborty T;

Wehland J; Heinz D W (Reprint)

CORPORATE SOURCE: UNIV FREIBURG, INST ORGAN CHEM & BIOCHEM, ALBERTSTR

21, D-79104 FREIBURG, GERMANY (Reprint); UNIV FREIBURG, INST ORGAN CHEM & BIOCHEM, D-79104 FREIBURG, GERMANY; GESELL BIOTECHNOL FORSCH MBH,

D-38124 BRAUNSCHWEIG, GERMANY; UNIV GIESSEN, INST

MED MIKROBIOL, D-35385 GIESSEN, GERMANY

COUNTRY OF AUTHOR: GERMANY

SOURCE: JO

JOURNAL OF MOLECULAR BIOLOGY, (17 OCT 1997) Vol.

273, No. 1, pp. 269-282.

Publisher: ACADEMIC PRESS LTD, 24-28 OVAL RD,

LONDON, ENGLAND NW1 7DX.

ISSN: 0022-2836.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT:

LIFE

LANGUAGE:

English 58

REFERENCE COUNT:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

The X-ray crystal structure of the phosphatidylinositol-AB specific phospholipase C (PT-PLC) from the human pathogen Listeria monocytogenes has been determined both in free form at 2.0 Angstrom resolution, and in complex with the competitive inhibitor myo-inositol at 2.6 Angstrom resolution. The structure was solved by a combination of molecular replacement using the structure of Bacillus cereus PI-PLC and single isomorphous replacement. The enzyme consists of a single (beta alpha)(8)-barrel domain with the active site located at the C-terminal side of the beta-barrel. Unlike other (beta alpha)(8)-barrels, the barrel in PI-PLC is open because it lacks hydrogen bonding interactions between beta-strands V and VI. myo-Inositol binds to the active site pocket by making specific hydrogen bonding interactions with a number of charged amino acid side-chains as well as a coplanar stacking interaction with a tyrosine residue. Despite a relatively low sequence identity of approximately 24%, the structure is highly homologous to that of B.cereus PI-PLC with an r.m.s. deviation for 228 common C-alpha positions of 1.46 Angstrom. Larger differences are found for loop regions that accommodate most of the numerous amino acid insertions and deletions. The active site pocket is also well conserved with only two amino acid replacements directly implicated in inositol binding. (C) 1997 Academic Press Limited.

L13 ANSWER 38 OF 47 ACCESSION NUMBER:

WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

1996-277797 [28] WPIDS

DOC. NO. CPI:

C1996-088243

TITLE:

Screening assay for inhibitors of nucleic acid polymerase(s) - using a polynucleotide template and

a polycationic polynucleotide-selective agent

immobilised on polymeric microbeads.

DERWENT CLASS:

B04 C07 D16

INVENTOR(S):

LAMARCO, K; STRULOVICI, B; WEI, P; WU, P

PATENT ASSIGNEE(S):

(TULA-N) TULARIK INC

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 9617084 A1 19960606 (199628) * EN 25

RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE

W: AT AU CA JP

AU 9643692 A 19960619 (199640)

21

US 5635349 A 19970603 (199728)

EP 793730 A1 19970910 (199741) EN

R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

ΑU	688190	В	19980305	(199820)		
JΡ	10510429	W	19981013	(199851)		31
CA	2205804	С	20001010	(200056)	EN	

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9617084 AU 9643692 US 5635349 EP 793730	A1 A A A	WO 1995-US15300 AU 1996-43692 US 1994-348797 EP 1995-942477	19951122 19951122 19941202 19951122
EP /93/30	AI	WO 1995-US15300	19951122
AU 688190	В	AU 1996-43692	19951122
JP 10510429	M	WO 1995-US15300	19951122
		JP 1996-518945	19951122
CA 2205804	С	CA 1995-2205804 WO 1995-US15300	19951122 19951122

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9643692 EP 793730 AU 688190	A Based on Al Based on B Previous Publ. Based on	WO 9617084 WO 9617084 AU 9643692 WO 9617084
JP 10510429	W Based on	WO 9617084
CA 2205804	C Based on	WO 9617084

PRIORITY APPLN. INFO: US 1994-348797 19941202

AN 1996-277797 [28] WPIDS

AB WO 9617084 A UPAB: 19960719

The following are claimed: (A) a method of identifying an inhibitor of a nucleic acid polymerase (NAP) activity comprising: (a) forming a mixt. of a nucleoside triphosphate (NTP), a polynucleotide template (PT), a pathogenic NAP and a candidate inhibitor of NAP activity, the NTP comprising a detectable label and the PT comprising a nucleotide sequence; (b) incubating the mixt. under conditions whereby, but for the presence of the candidate inhibitor, the polymerase transcribes the PT by catalysing the polymerisation of the NTP into a polynucleotide comprising a nucleotide sequence complementary to that of the PT; (c) contacting the mixt. with a polycationic polynucleotide-selective agent immobilised on polymeric microbeads; (d) incubating the mixt. in the presence of the solid substrate to selectively bind the polynucleotide to the polynucleotide selective agent; (e) sepg. the polymeric microbeads from the mixt.; (f) washing the microbeads free of the NTP, and (g) detecting the presence or absence of the label on the microbeads, whereby the absence of the label on the microbeads indicates that the candidate inhibitor is an inhibitor of a pathogenic NAP activity; (B) a membrane filtration appts. comprising a tube having a fluid passage comprising a reservoir portion, a first filter and a second filter, each of the first and second filters extending transversely across the passage, the first filter positioned between the reservoir portion of the passage and the second filter, the first filter being changeable from water-impermeable to

water-permeable by contact with an organic solvent, and the second filter being water-permeable, hydrophilic, capable of permitting the passage of 90% by vol. of free water from the reservoir portion of the passage while retaining 90% by vol. of particles with a size of 20-200 mu dia. in the reservoir portion of the passage and having a max. pore size < 200 mu dia.

USE - The method provides for the high-throughput screening for specific inhibitors of pathogenic NAP activity. The inhibitors can be used as reagents in a wide variety of in vitro and cellular applications, in plant and field crops, pesticides, fungicides and in animal and human trials for diagnostic and therapeutic applications. They can also be used in studies on in vitro gene transcription systems. Dwq.0/2

5635349 A UPAB: 19970709 ABEQ US

A method of identifying an inhibitor of a nucleic acid polymerase activity, the method comprises: forming a mixture of nucleoside triphosphates, a polynucleotide template, a pathogenic nucleic acid polymerase and a candidate inhibitor of nucleic acid polymerase activity, at least one of the nucleoside triphosphate comprising a detectable label and the polynucleotide template comprising a nucleotide sequence;

incubating the mixture under conditions whereby, but for the presence of the candidate inhibitor, the polymerase transcribes the polynucleotide template by catalysing the polymerization of the nucleoside triphosphates into a polynucleotide comprising a nucleotide sequence complementary to that of the polynucleotide template; (b) contacting the mixture with a polycationic, non-sequence-specific, polynucleotide-selective agent immobilised on polymeric microbeads; (c) incubating the mixture in the presence of the polymeric microbeads under conditions to selectively bind the polynucleotide to the polynucleotide-selective agent; (d) separating the polymeric microbeads from the mixture by membrane filtration; (e) washing the polymeric microbeads free of the nucleoside triphosphate; and (f) detecting the presence or absence of the label on the polymeric microbeads.

The absence of the label on the polymeric microbeads indicates that the candidate inhibitor of polymerase activity is an inhibitor of a pathogenic nucleic acid polymerase activity. Dwg.0/2

DUPLICATE 2 L13 ANSWER 39 OF 47 MEDLINE

ACCESSION NUMBER:

96433078 MEDLINE

PubMed ID: 8836106 DOCUMENT NUMBER: 96433078

Crystal structures of Toxoplasma gondii TITLE:

HGXPRTase reveal the catalytic role of a long

flexible loop.

Comment in: Nat Struct Biol. 1996 Oct;3(10):813-4 COMMENT:

Schumacher M A; Carter D; Ross D S; Ullman B; Brennan AUTHOR:

Department of Biochemistry and Molecular Biology, CORPORATE SOURCE:

Oregon Health Sciences University, Portland

97201-3098, USA.

NATURE STRUCTURAL BIOLOGY, (1996 Oct) 3 (10) 881-7. SOURCE:

Journal code: B98; 9421566. ISSN: 1072-8368.

United States PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

> Shears 308-4994 Searcher :

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199611

ENTRY DATE: Entered STN: 19961219

Last Updated on STN: 19961219 Entered Medline: 19961114

AB Crystal structures of substrate-free and

XMP-soaked hypoxanthine-guanine-xanthine phosphoribosyltransferase (HGXPRTase) of the opportunistic pathogen Toxoplasma gondii have been determined to 2.4 and 2.9 A resolution, respectively. HGXPRTase displays the conserved PRTase fold. In the structure of the enzyme bound to its product, a long flexible loop (residues 115-126) is located away from the active site. Comparison to the substrate-free structure reveals a striking relocation of the loop, which is poised to cover the catalytic pocket, thus providing a mechanism by which the HG(X)PRTases shield their oxocarbonium transition states from nucleophilic attack by the bulk solvent. The conserved Ser 117-Tyr 118 dipeptide within the loop is brought to the active site, completing the ensemble of catalytic residues.

L13 ANSWER 40 OF 47 MEDLINE DUPLICATE 3

ACCESSION NUMBER: 95171111 MEDLINE

DOCUMENT NUMBER: 95171111 PubMed ID: 7866745
TITLE: Crystal structure of scytalone

dehydratase -- a disease determinant of the

rice pathogen, Magnaporthe grisea.

AUTHOR: Lundqvist T; Rice J; Hodge C N; Basarab G S; Pierce

J; Lindqvist Y

CORPORATE SOURCE: Department of Molecular Biology, Swedish University

of Agricultural Sciences Uppsala Biomedical Center.

SOURCE: STRUCTURE, (1994 Oct 15) 2 (10) 937-44.

Journal code: B31; 9418985. ISSN: 0969-2126.

PUB. COUNTRY: ENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199503

ENTRY DATE: Entered STN: 19950407

Last Updated on STN: 19990129 Entered Medline: 19950329

BACKGROUND: Rice blast is caused by the pathogenic AB fungus, -Magnaporthe grisea. Non-pathogenic mutants have been identified that lack enzymes in the biosynthetic pathway of dihydroxynapthalene-derived melanin. These enzymes are therefore prime targets for fungicides designed to control rice blast disease. One of the enzymes identified by genetic analysis as a disease determinant is scytalone dehydratase. RESULTS: The three-dimensional structure of scytalone dehydratase in complex with a competitive inhibitor has been determined at 2.9 A resolution. A novel fold, a cone-shaped alpha + beta barrel, is adopted by the monomer in this trimeric protein, burying the hydrophobic active site in its interior. The interactions of the inhibitor with the protein side chains have been identified. The similarity of the inhibitor to the substrate and the side chains involved in binding afford some insights into possible catalytic mechanisms. CONCLUSIONS: These results provide a first look into the structure and catalytic residues of a non-metal dehydratase, a large class of hitherto structurally uncharacterized enzymes. It is envisaged that a

detailed structural description of scytalone dehydratase will assist in the design of new inhibitors for controlling rice blast disease.

L13 ANSWER 41 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

1993-303652 [38] WPIDS

DOC. NO. NON-CPI: DOC. NO. CPI:

N1993-233444 C1993-135305

TITLE:

Detection of antibodies to

pathogenic virus - by reaction of

micro-capillary spot samples with enzyme-labelled viral antigen, used for mass screening mink for

Aleutian disease. B04 D16 J04 S03

DERWENT CLASS: INVENTOR(S):

MIROSHNICHENKO, S M; PEREMYSLOV, V V; TARANIN, A V

PATENT ASSIGNEE(S):

(TARA-I) TARANIN A V

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
WO 9318403	A2	19930916	(199338)*	RU	12
W: DK US	-		44.0044.01		
DK 9301288	Α	19940110	(199412)		
RU 2074393	C1	19970227	(199740)		7

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9318403 DK 9301288	A2 A	WO 1993-RU33 WO 1993-RU33 DK 1993-1288	19930210 19930210 19931115
RU 2074393	C1	SU 1992-5032100	19920313

PRIORITY APPLN. INFO: SU 1992-5032100 19920313

AN 1993-303652 [38] WPIDS

9318403 A UPAB: 19931123 AB

> Detection of antibodies to a pathogenic virus is effected by depositing samples on a protein-binding porous solid support in the form of microcapillary spots; drying the samples; directly reacting any antibodies in the samples with an enzyme-labelled viral antigen; and detecting any label by colour reaction with a substrate.

The support is pref. a sheet of nitrocellulose with ruled lines forming an array of squares for receiving the samples. Each square is numbered or lettered to identify the sample. Blood samples are diluted 2- to 20-fold with phosphate-buffers saline. The enzyme is a peroxidase, phosphatase or galactosidase and is linked to the antigen directly or via suitable ligands, esp. biotin and streptavidin.

USE/ADVANTAGE - The method is highly sensitive (e.g. with a detection limit of 0.01-0.07 mcg/ml), require only small samples, does not require complex equipment or highly qualified personnel, and can be used to test large nos. of biological fluid samples of any kind in a short time (e.g. 2.25 hr. for 100-1000 samples).

Dwg.0/0

L13 ANSWER 42 OF 47 MEDLINE DUPLICATE 4

ACCESSION NUMBER: 93193345 MEDLINE

DOCUMENT NUMBER: 93193345 PubMed ID: 8448924

TITLE: Inhibition of mitochondrial respiration by

furancarboxylic acid accumulated in uremic serum in

its albumin-bound and non-dialyzable form.

AUTHOR: Niwa T; Aiuchi T; Nakaya K; Emoto Y; Miyazaki T;

Maeda K

CORPORATE SOURCE: Department of Internal Medicine, Nagoya University

Branch Hospital, Japan.

SOURCE: CLINICAL NEPHROLOGY, (1993 Feb) 39 (2) 92-6.

Journal code: DEY; 0364441. ISSN: 0301-0430.

PUB. COUNTRY: GERMANY: Germany, Federal Republic of

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199304

ENTRY DATE: Entered STN: 19930423

Last Updated on STN: 19930423 Entered Medline: 19930409

3-carboxy-4-methyl-5-propyl-2-furanpropionic acid (CMPF) accumulates AΒ markedly in uremic serum in its albumin-bound form. To determine if CMPF can be removed by newly developed dialyzers with high-flux membranes which are permeable to low-molecular-weight proteins, such as beta 2-microglobulin (beta 2-MG), serum levels of CMPF were determined before and after hemodialysis using these high-flux membrane dialyzers. In addition, to determine the pathogenic role of CMPF in uremic patients, its cellular toxicity due to its effect on mitochondrial respiration was studied. The reduction rates of CMPF by hemodialysis using the dialyzers ranged from -17% to -24%, demonstrating the nondialyzability of CMPF due to its strong albumin-binding, while those of beta 2-MG ranged from 11% to 43%. CMPF inhibited ADP-stimulated oxidation of NADH-linked substrates in isolated mitochondria dose-dependently regardless of the presence of serum albumin. This inhibition was observed even at a concentration of 0.2 mM, which is comparable to the serum levels of CMPF in the hemodialysis patients. In conclusion CMPF which cannot be removed even by high-flux membrane dialyzers, is a strong inhibitor of mitochondrial respiration, and novel purification methods to remove CMPF from the blood of uremic patients should be developed.

L13 ANSWER 43 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1992-080211 [10] WPIDS

ACCESSION NUMBER: 1992-080211 [10]
DOC. NO. NON-CPI: N1992-060054

DOC. NO. CPI: N1992 000034

TITLE: New adhesion receptors for pathogenic and

opportunistic microorganisms - useful as vaccines and for diagnosis, treatment and prevention of pathogenic and opportunistic infections e.g.

salmonella.

DERWENT CLASS: B04 D16 S03

INVENTOR(S): KRIVAN, H C; SAMUEL, J E

PATENT ASSIGNEE(S): (ANTE-N) ANTEX BIOLOGICS INC; (BIOC-N) BIOCARB INC;

(MICR-N) MICROCARB INC

COUNTRY COUNT: 17

PATENT INFORMATION:

PAT	CENT NO	KIND	DATE	WEEK	LA	PG
WO	9202817	-	19920)220 (199	210)*	67
	RW: AT BE	CH I	E DK	ES FR GE	GR IT LU	NL SE
	W: CA JP					
ΕP	553113	A1	19930	0804 (199	331) EN	67
	R: AT CH	DE I	K ES	FR GB IT	LI NL SE	
JΡ	06501383	M	19940	217 (199	(412)	19
EP	553113	A4	19940	330 (199	530)	
US	5696000	Α	19971	209 (199	804)	20
ΕP	553113	В1	19981	125 (199	9851) EN	
	R: AT CH	DE [OK ES	FR GB IT	LI NL SE	
DE	69130536	E	19990	107 (199	907)	
ES	2127198	Т3	19990)416 (199	922)	
					018) EN	

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 553113	A1	EP 1991-916508 WO 1991-US5179	19910729 19910729
JP 06501383	W	JP 1991-515061 WO 1991-US5179	19910729 19910729
EP 553113	A4	EP 1991-916508	10000000
US 5696000	A Div ex Cont of	US 1990-562002 US 1993-78660	19900802 19930621
		US 1994-275702	19940718 19910729
EP 553113	B1	EP 1991-916508 WO 1991-US5179	19910729
DE 69130536	E	DE 1991-630536	19910729
		EP 1991-916508	19910729
		WO 1991-US5179	19910729
ES 2127198	Т3	EP 1991-916508	19910729
CA 2095642	С	CA 1991-2095642	19910729
		WO 1991-US5179	19910729

FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 553113 JP 06501383 EP 553113 DE 69130536 ES 2127198	Al Based on W Based on Bl Based on E Based on Based on T3 Based on	WO 9202817 WO 9202817 WO 9202817 EP 553113 WO 9202817 EP 553113
CA 2095642	C Based on	WO 9202817

PRIORITY APPLN. INFO: US 1990-562002 19900802; US 1993-78660 19930621; US 1994-275702 19940718

1992-080211 [10] WPIDS AN

WO 9202817 A UPAB: 19970716 AΒ

A receptor (I) comprises a substantially pure compound e.g. GalB1-4GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x(R), GalB1-4GlcNAcB1-3GalB1-4GlcB1-1-x(R), GalB1-4GlcNAcB1-3GalB1-4Glc, GalB1-3G1cNAcB1-3GalB1-4Glc, GlcNAbi-3GalB1-4Glc, GalB1-4GlNAc-3Gal

> 308-4994 Shears Searcher :

and GalB1-3-GlcNAcB1-3Gal, where x is sphingosine, hydroxylated sphingosine or saturated sphingosine, and R is H or N-acyl fatty acid derivative of x such that x(R) is a ceramide.

Also new are (1) a compsn. comprising (I) attached to an insoluble or soluble substrate; (2) a method for detecting microorganisms in a sample comprising:— (a) contacting the sample with (I) for a period of time and under conditions sufficient for the receptors to bind the microorganisms; and (b) assaying for complexes formed; (3) a diagnostic kit for the detection of pathogenic or opportunistic microorganisms comprising the compsn. of (1) and means for detecting or measuring the formation of complexes.

USE/ADVANTAGE - (I) can be used in various forms to detect microorganisms, remove them from liquids and treat infections caused by them. It inhibits their adhesion to mammalian cells if suspended in a liquid which is physiologically compatible with the cells. The adhesion protein has diagnostic and therapeutic applications and is particularly useful as a vaccine. @(67pp Dwg.No.0/4

ABEO EP 553113 A UPAB: 19931118

A receptor (I) comprises a substantially pure compound e.g. GalB1-4GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-1-x (R), GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-3GlcNAcB1-3GalB1-4Glc, GalB1-4Glc, GalB1-4Glc, GalB1-4Glc, GalB1-4Glc, GalB1-3GlcNAcB1-3Gal, where x is sphingosine, hydroxylated sphingosine or saturated sphingosine, and R is H or N-acyl fatty acid derivative of x such that x(R) is a ceramide.

Also new are (1) a compsn. comprising (I) attached to an insoluble or soluble substrate; (2) a method for detecting microorganisms in a sample comprising:— (a) contacting the sample with (I) for a period of time and under conditions sufficient for the receptors to bind the microorganisms; and (b) assaying for complexes formed; (3) a diagnostic kit for the detection of pathogenic or opportunistic microorganisms comprising the compsn. of (1) and means for detecting or measuring the formation of complexes.

USE/ADVANTAGE - (I) can be used in various forms to detect microorganisms, remove them from liquids and treat infections caused by them. It inhibits their adhesion to mammalian cells if suspended in a liquid which is physiologically compatible with the cells. The adhesion protein has diagnostic and therapeutic applications and is particularly useful as a vaccine.

ABEQ US 5696000 A UPAB: 19980126

A receptor (I) comprises a substantially pure compound e.g. GalB1-4GlcNAcB1-3GalB1-4GlcB1-l-x (R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-l-x(R), GalB1-3GlcNAcB1-3GalB1-4GlcB1-l-x(R), GalB1-4GlcNAcB1-3GalB1-4Glc, GalB1-3GlcNAcB1-3GalB1-4Glc, GlcNAb1-3GalB1-4Glc, GalB1-4GlNAc-3Gal and GalB1-3-GlcNAcB1-3Gal, where x is sphingosine, hydroxylated sphingosine or saturated sphingosine, and R is H or N-acyl fatty acid derivative of x such that x(R) is a ceramide.

Also new are (1) a compsn. comprising (I) attached to an insoluble or soluble substrate; (2) a method for detecting microorganisms in a sample comprising:— (a) contacting the sample with (I) for a period of time and under conditions sufficient for the receptors to bind the microorganisms; and (b) assaying for complexes formed; (3) a diagnostic kit for the detection of pathogenic or opportunistic microorganisms comprising the compsn. of (1) and means for detecting or measuring the formation of complexes.

USE/ADVANTAGE - (I) can be used in various forms to detect microorganisms, remove them from liquids and treat infections caused by them. It inhibits their adhesion to mammalian cells if suspended in a liquid which is physiologically compatible with the cells. The adhesion protein has diagnostic and therapeutic applications and is particularly useful as a vaccine. Dwg.0/4b

L13 ANSWER 44 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD ACCESSION NUMBER: 1992-049529 [07] WPIDS 1992-390495 [48]; 1992-390496 [48] CROSS REFERENCE: DOC. NO. NON-CPI: N1992-037843 C1992-022044 DOC. NO. CPI: Appts. for immunoassay of bacterial TITLE: lipo-polysaccharide - has substrate coated with polymyxin in place of capture antibody, esp. for detecting Salmonella in food. A88 A96 B04 B07 C07 D13 D16 J01 J04 P34 S03 DERWENT CLASS: BLAIS, B W; YAMAZAKI, H INVENTOR(S):

(BLAI-I) BLAIS B W; (YAMA-I) YAMAZAKI H PATENT ASSIGNEE(S):

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
CA 2017093 JP 04270965 US 5510242	Α	19911118 19920928 19960423			23 17

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
CA 2017093 JP 04270965 US 5510242	A A A Cont of	CA 1990-2017093 JP 1991-113467 US 1991-697683 US 1993-87013	19900518 19910517 19910509 19930707

PRIORITY APPLN. INFO: CA 1990-2017093 19900518; CA 1991-2037726 19910307; CA 1991-2037727 19910307

1992-049529 [07] ΑN WPIDS

1992-390495 [48]; 1992-390496 [48] CR

2017093 A UPAB: 19931006 AB

The appts. comprises a substrate to which polymyxin (I) is adhered and to which the sample is applied. Also new is detection of bacterial LKPS using this appts...

The assay can pref. be on enzyme, radio or fluorescent immunoassay and the substrate is plastic (specifically polystyrene, polycarbonate, polymethacrylate or PVC); woven or non-woven cloth (pref. rayon-polyester) or paper. (I) is polymyxin B, B1, B2, D1, D2 or E.

USE/ADVANTAGE - (I) binds with LPS of all Gram-negative bacteria, and is much cheaper than captive antibodies used previously. The new appts. does not need to be refrigerated; provides a rapid test (within 24 hrs.) which can be made semi-quantitative; and can detect as few as 1-10 Salmonella cells per g of food. Unlike antibodies, (I) has consistent

> Shears 308-4994 Searcher :

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batch-to-batch quality. The method is used to
detect bacterial pathogens esp. in food.
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5510242 A UPAB: 19960604 ABEO US

A process for the detection of lipopolysaccharide present on the cell walls of a target Gram negative bacteria which process comprises the steps of:

- a) heating a sample of bacteria in a detergent solution to extract lipopolysaccharide antigens from said cell walls of said Gram negative bacteria;
- b) contacting said solution with a device consisting essentially of in combination, a macroporous, hydrophobic material as a substrate, said substrate comprising a cloth composed of hydrophobic synthetic polymeric fibers, selected from the group consisting of polyester, polypropylene, and nylon and blends thereof with rayon which are either woven or non-woven into a physically-structurally-stable cloth of more than about 200 mum thickness, such that the pores exceed about 20 mum in diameter, said substrate being adapted to receive a sample to be tested, and, bound by simple adsorption to said substrate, a polymyxin;

c) washing said device; and

d) detecting the presence of said predetermined lipopolysaccharide by contacting an antibody indicator conjugate specifically binding said lipopolysaccharide to said device. Dwg.0/7

L13 ANSWER 45 OF 47 WPIDS COPYRIGHT 2002 ACCESSION NUMBER:

DERWENT INFORMATION LTD

1990-209784 [27] WPIDS

DOC. NO. NON-CPI:

N1990-163010

DOC. NO. CPI: TITLE:

C1990-090657 Amplified DNA capture and detection - by

incorporating ligand and contacting DNA with

substrate having immobilised binding reagent for ligand.

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

FOOTE, S J; KEMP, D J; PETERSON, M G; SAMARAS, N;

SMITH, D

PATENT ASSIGNEE(S):

(AMRA-N) AMRAD CORP LTD; (AMRA) AMERICAN STANDARD

INC

COUNTRY COUNT:

18

PATENT INFORMATION:

PAT	ENT NO	KIND	DATE	WEEK	LA	PG
	9006374	Δ	19900614	(199027)	*	
WO.	RW: AT BE					
	W: AU DK					
CA	2004990	Α	19900609	(199034)		
	8946637					
EΡ	447464	A	19910925	(199139)		
			DE ES FR			SE
			19910807			
	9102169					
JΡ	04502251					25
ΑU	633036	В	19930121			
EΡ	447464	A4	19920603	(199522)		

US	5536648	A 19960716 (199634)	23
ΕP	447464	B1 19980805 (199835) EN	
	R: AT BE	CH DE ES FR GB IT LI LU NL	SE
DE	68928769	E 19980910 (199842)	
ES	2121750	T3 19981216 (199906)	
.TP	3068848	B2 20000724 (200040)	20

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 447464	A	EP 1990-900722	19891208
JP 04502251	W	WO 1989-AU526	19891208
		JP 1990-500923	19891208
AU 633036	В .	AU 1989-46637	19891208
EP 447464	A4	EP 1990-900722	
US 5536648	A Cont of	WO 1989-AU526	19891208
	Cont of	US 1991-689287	19910801
		US 1994-229056	19940418
EP 447464	B1	WO 1989-AU526	19891208
		EP 1990-900722	19891208
DE 68928769	E	DE 1989-628769	19891208
,,,,		WO 1989-AU526	19891208
		EP 1990-900722	19891208
ES 2121750	Т3	EP 1990-900722	19891208
JP 3068848	В2	WO 1989-AU526	19891208
		JP 1990-500923	19891208

FILING DETAILS:

PATENT NO	KIND	PATENT NO
JP 04502251 AU 633036	W Based on B Previous Publ. Based on	WO 9006374 AU 8946637 WO 9006374
EP 447464 DE 68928769	B1 Based on E Based on	WO 9006374 EP 447464
ES 2121750 JP 3068848	Based on T3 Based on B2 Previous Publ.	WO 9006374 EP 447464 JP 04502251
	Based on	W∩ 9006374

PRIORITY APPLN. INFO: AU 1988-1889 19881209; AU 1989-5080 19890704

AN 1990-209784 [27] WPIDS

AB WO 9006374 A UPAB: 19950524

Capturing amplified target DNA on a solid **substrate** is claimed comprising incorporating a first ligand into the DNA by a polymerase chain reaction (PCR) using a set of primers where one of the primers bears the ligand and contacting the treated DNA with a solid **substrate** having an immobilised **binding** reagent for the ligand.

The **binding** reagent may be a DNA **binding** protein, e.g. glutathione-5-transterase (GST)-GCN4 or Tyr R. Detection of the captured DNA may be with a second ligand, e.g. biotin and a detection reagent, e.g. avidin/peroxidase.

USE/ADVANTAGE - The single or multi-step amplified DNA assay provides a very sensitive, specific and rapid method for

detecting specific DNA segments. It can be used for screening for genetic disorders or pathogens, e.g. HIV. @(64pp Dwg.No.1/15)@
1/15@

ABEQ JP 04502251 W UPAB: 19930928

Capturing amplified targart DNA on a solid **substrate** is claimed comJP4502251A - Wprising incorporating a first ligand into the DNA by a polymerase chain reaction (PCR) using a set of primers where one of the primers bears the ligand and contacting the DNA with a solid **substrate** having an immobilised **binding** reagent for the ligand.

The **binding** reagent may be a DNA **binding** protein, e.g. glutathione-5-transferase (GST)-GCN4 or Tyr R. Detection of the captured DNA may be with a second ligand, e.g. biotin and a detection reagent, e.g. avidin/peroxidase.

USE/ADVANTAGE - The single or multi-step amplified DNA assay provides a very sensitive, specific and rapid method for detecting specific DNA segments. It can be used for screening for genetic disorders or pathogens, e.g. HIV.

ABEQ US 5536648 A UPAB: 19960829

A method for capturing target DNA on a solid substrate comprising: a) amplifying said target DNA by a polymerase chain reaction with a pair of oligonucleotide primers which are complementary to said target DNA wherein one of the primers comprises a nucleotide sequence which is a ligand for a double stranded DNA-binding protein when said nucleotide sequence is incorporated into said amplified target DNA; and b) contacting said amplified target DNA with a double stranded DNA-binding protein immobilized on a solid substrate.

Dwg.0/15

L13 ANSWER 46 OF 47 MEDLINE

ACCESSION NUMBER: 85090449 MEDLINE

DOCUMENT NUMBER: 85090449 PubMed ID: 2578226

TITLE: Fluorogenic substrate detection of viable intracellular and extracellular

pathogenic protozoa.

AUTHOR: Jackson P R; Pappas M G; Hansen B D SOURCE: SCIENCE, (1985 Jan 25) 227 (4685) 435-8.

Journal code: UJ7; 0404511. ISSN: 0036-8075.

PUB. COUNTRY: United States

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198502

ENTRY DATE: Entered STN: 19900320

Last Updated on STN: 19900320 Entered Medline: 19850221

Viable Leishmania promastigotes and amastigotes were detected by epifluorescence microscopy with fluorescein diacetate being used to mark living parasites and the nucleic acid-binding compound ethidium bromide to stain dead cells. This procedure is superior to other assays because it is faster and detects viable intracellular as well as extracellular Leishmania. Furthermore, destruction of intracellular pathogens by macrophages is more accurately determined with fluorescein diacetate than with other stains. The procedure may have applications in programs

to develop drugs and vaccines against protozoa responsible for human and animal disease.

L13 ANSWER 47 OF 47 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1984-258210 [42] WPIDS

DOC. NO. NON-CPI: N1984-192939 DOC. NO. CPI: C1984-109158

TITLE: Enzymatic channelling binding immuno

assay - for measurement of analytes, such as

poly-nucleotide(s).

DERWENT CLASS: A89 B04 D16 J04 S03

INVENTOR(S): DINELLO, R; GIBBONS, I; ULLMAN, E F

PATENT ASSIGNEE(S): (SYNT) SYVA CO

COUNTRY COUNT: 16

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA	PG
EP 122059		17 (198442)		36
AU 8425561		20 (198445)	
JP 59178362 US 4687735				
CA 1230289 EP 122059				
R: AT BE	CH DE FR G	B IT LI LU	NL SE	
DE 3476834 JP 06043997				11

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 122059	A	EP 1984-301676	19840313
JP 59178362	A	JP 1984-46614	19840313
US 4687735	A	US 1983-474906	19830314
TP 06043997	B2	JP 1984-46614	19840313

FILING DETAILS:

PATENT NO	KIND	PATENT NO
TP 06043997	B2 Based on	JP 59178362

PRIORITY APPLN. INFO: US 1983-474906 19830314

AN 1984-258210 [42] WPIDS

AB EP 122059 A UPAB: 19930925

A channelling assay method for detecting the presence of an analyte, which is a member of a specific binding pair comprising first and second binding members (FBM and SBM resp.), is claimed. The method employs two systems: (A) a channelling signal producing system comprising (i) an enzyme (E1)-SBM conjugate; (ii) a substrate for E1; (iii) a final reactant which is either another enzyme (E2) related to E1 by the substrate of one enzyme being the prod. of the other, or a cpd. which reacts with the prod. of E1 reaction; and (B) a linking system causing polymerisation of the final reactant and incorporation of E1 within the polymer as a function of the

binding of E1-SBM conjugate to FMB so as to form a polymeric channelling aggregate.

The process is performed in an aq. assay medium by combining the sample, E1-SBM conjugate (and FBM when the analyte is a SBM) and the linking system to form the aggregate, through which is then channelled the remaining components of system A, to form a final prod., which causes a change in a detectable signal. The signal is then compared with that observed in a medium contg. a known amount of analyte.

Pref. the FBM is a ligand and the SBM is a receptor, pref. an antibody. The enzymes are oxidoreductases.

USE - The assay may be used to detect analytes in physiological fluids, cellular materials, etc. or it may be employed in monitoring contaminants in water and chemical processing etc. The analyte is pref. a ligand such as a drug, macromolecule, protein or polynucleotide. Suitable ligands are described in US4233402.

ABEO EP 122059 B UPAB: 19930925

A channelling assay method for detecting the presence of an analyte in a sample suspected of containing said analyte, when said analyte is a member of a specific binding pair consisting of first and second binding members ("FBM" and "SBM", respectively), characterised by said method employing a channelling signal producing system comprising an enzyme, enzyme(+), conjugated to an SBM to provide enzyme (+)-SBM conjugate, an enzyme substrate and a Final Reactant, wherein said Final Reactant is (1) another enzyme, enzyme(x), where the two enzymes are related by the substrate of one enzyme being the product of the other enzyme, or (2) a compound which reacts with the product of enzyme(+), wherein the reactions of said enzyme(+) in conjunction with enzyme(x) or said compound results in a change in an observable signal in relation to the amount of analyte in said sample; and linking system providing for the polymerisation of said Final Reactant and the incorporation of enzyme(+) within the polymer as a function of the binding of enzyme(+)-SBM conjugate to FBM so as to form a polymeric channelling aggregate; said method comprising: combining in an aqueous assay medium; (a) said sample; (b) said enzyme(+)-SBM conjugate, and FBM when said analayte is a SBM, so as to form a complex between said FBM and said enzyme(+)-SBM conjugate in relation to the amount of analyte present; (c) any members of said linking system, whereby is formed a polymeric channelling aggregate of said Final Reactant and a complex of enzyme(+)-SBM conjugate with FBM; and (d) remaining members of a signal producing system, whereby a Final Product is produced as a result of channelling of said members of said signal producing system in said polymeric channelling aggregate, which results in a change in detectable signal; and comparing said detectable signal to the detectable signal observed in an assay medium having a known amount of analyte.

ABEQ US 4687735 A UPAB: 19930925

Detection of an analyte which is a member of a specific binding pair FBM-SBM uses a channeling signal producing system comprising an enzyme, designated enzyme+, conjugated to an SBM to provide enzyme+ -SBM conjugate, an enzyme substrate and a Final Reactant. The Final Reactant is (1) another enzyme designated enzyme, such that the prod. of one enzyme is the substrate of the other or (2) a cpd. reactive with the prod. of enzyme+, in either case reaction producing an observable signal

related to the amt. of analyte in the test sample. A linking system provides for polymerisation of the Final Reactant and incorporation of enzyme+ within the polymer chain as a function of the binding of enzyme+ -SBM conjugate to FBM so as to form a polymeric channeling aggregate.

USE/ADVANTAGE - In determn. of pathogens, etc. with increased sensitivity.

> LUS' ENTERED AT 15:42:13 ON 06 MAR 2002 Claim 46

134 S L1 AND KIT

10 S L14 AND SUBSTRATE L15

8 S L15 NOT L9 L16

L14

L16 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:904467 CAPLUS

DOCUMENT NUMBER: 136:32654

Primer extension using a mixture of labeled and TITLE:

unlabeled nucleotides for detection of mutation

in genes

Xu, Hua INVENTOR(S):

Dna Sciences, Inc., USA PATENT ASSIGNEE(S): PCT Int. Appl., 63 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE					APPLICATION NO. DATE											
WO.	WO 2001094546			 A	- - 2	20011213			WO 2001-US179			28	28 20010531			
	W:	AE,	AG,	AL,	AM,	AT,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,
		CH,	CN,	CR,	CU,	CZ,	CZ,	DE,	DE,	DK,	DK,	DM,	DZ,	EE,	EE,	ES,
		FI,	FI.	GB,	GD,	GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	ΚE,
		KG,	KP,	KR,	KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,
		MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,
		SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	YU,	ZA,	ZW,
						KZ,										
	RW:	GH,	GM,	KE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	ΤZ,	UG,	ZW,	ΑT,	BE,	CH,
		CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PΤ,	SE,
		TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	SN,	TD,
		TG														
RITY	APP	LN.	INFO	. :					US 2	000-	5857	68	Α2	2000	0602	

PRIORIT The present invention provides methods, compns. and kits for detg. the identity of a nucleotide at a variant site in a nucleic acid of interest, including point mutations and single nucleotide polymorphism. The methods utilize one or more nucleotides, each nucleotide being a mixt. of labeled and unlabeled forms, to generate labeled extension products that are characteristic of the nucleotide at the variant site in the nucleic acid of interest. In addn. to their utility in detecting and analyzing point mutations and SNPs, the methods and kits of the invention have utility in a variety of other applications in which specific nucleic acid sequence information is of value, including detection of pathogens, paternity disputes, prenatal testing and forensic anal.

L16 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2002 ACS

2001:886567 CAPLUS ACCESSION NUMBER: 136:32642 DOCUMENT NUMBER: Determination of DNA sequence variations through TITLE: primer extension and uses in genotyping Xu, Hua; Glazer, Alexander N. INVENTOR(S): Dna Sciences, Inc., USA PATENT ASSIGNEE(S): PCT Int. Appl., 54 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: APPLICATION NO. DATE PATENT NO. KIND DATE ______ ____ _____ WO 2001-US18023 20010531 20011206 WO 2001092583 A1 AE, AG, AL, AM, AT, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EE, EE, ES, FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 2000-586125 A2 20000602 PRIORITY APPLN. INFO.: The present invention provides methods and kits for detg. the identity of a nucleotide at a variant site in a target nucleic acid of interest, including, for example, point mutations and single nucleotide polymorphisms. The methods involve conducting template-dependent extension reactions in the presence of a mixt. of nucleotides including labeled extendible nucleotide and labeled non-extendible nucleotide that are selected to be complementary to the nucleotides that potentially occupy the variant site. The particular labeled nucleotide incorporated into the extension products is characteristic of the nucleotide at the variant site. The methods can be used in conducting genotyping analyses and can be performed in multiplexing formats. In addn. to their utility in analyzing point mutations and single nucleotide polymorphisms, the methods and kits of the invention have utility in a variety of other applications in which specific nucleotide sequence information is of value, including, for example, paternity disputes, prenatal testing, forensic anal. and detection of pathogens. THERE ARE 1 CITED REFERENCES AVAILABLE FOR REFERENCE COUNT: 1 THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L16 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2002 ACS 2001:798475 CAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 135:353697 Detecting gene mutations or polymorphisms via TITLE: the proofreading activity of polymerases and uses thereof

Searcher: Shears 308-4994

Xu, Hua; Mathies, Richard A.

DNA Sciences, Inc., USA

INVENTOR(S):

PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 82 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT	NO.		KIN	1D	DATE			A	PPLI	CATI	ои ис	o. I	DATE		
	WO 2001	08163	1	A1	_	2001	1101		W	20	01-U	3131	36	2001	0424	
	W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,
		CN,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	KP,	KR,	ΚŻ,	LC,	LK,
		LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,
		PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,
				US,	UZ,	VN,	YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	KZ,	MD,	RU,
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	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	52,	TZ,	TIT	ΔW,	MI,	םם, סידים	CR,
		CY,	DE,	DK,	ES,	CG,	EK,	GB,	GK,	CM	CW,	ыU,	MD,	NE,	CNI	TD
		TK, TG	Br,	во,	CF,	CG,	CI,	CM,	GA,	GIV,	GW,	mы,	PIK,	ME,	DIV,	10,
DDTC	RITY APP		NEO						tis 21	000-	5582	4.5	A2 :	2000	0425	
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	of cert	ain D	NA p	olyn	nera	ses :	in c	onju:	ncti	on w	ith:	sele	cted	det	ecti	on
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L16 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2001:472904 CAPLUS

THE RE FORMAT

DOCUMENT NUMBER:

135:73344

TITLE:

RNA polymerases from bacteriophage .phi.6-.phi.14 and their use for

primer-independent RNA synthesis Makeyev, Eugeny; Bamford, Dennis

INVENTOR(S): PATENT ASSIGNEE(S):

Finland

SOURCE:

PCT Int. Appl., 63 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

Shears 308-4994 Searcher :

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APPLICATION NO.
     PATENT NO.
                      KIND
                             DATE
                      ____
     WO 2001046396
                             20010628
                                           WO 2000-FI1135
                                                             20001221
                      A1
         W: AE, AG, AL, AM, AT, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA,
             CH, CN, CR, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EE, EE, ES,
             FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK,
             MN, MW, MX, MZ, NO, NZ, PL
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH,
             CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
             TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD,
             TG
                                         FI 1999-2751
                                                          A. 19991221
PRIORITY APPLN. INFO.:
     A polymerase originating from a dsRNA virus catalyzes RNA synthesis
     using ssRNA, dsRNA, ssDNA, or dsDNA templates is disclosed. Such a
     polymerase can be purified from a dsRNA virus, and a protein having
     the amino acid sequence of such a polymerase is useful in methods
     and kits for in vitro RNA synthesis. A polymerase of the
     invention is processive, has very high RNA-polymn. rate and does not
     require primer for the initiation of RNA synthesis, although it is
     also able to initiate RNA synthesis in the presence of a primer.
     Expression and purifn. of recombinant P2 polymerase of bacteriophage
     .phi.6 is described. Nucleotide sequence of the P2 gene of
     bacteriophage .phi.6 and amino acid sequence of the encoded RNA
     polymerase are provided. RNA synthesis by the .phi.6 RNA polymerase
     in the reaction mixts. programmed with dsRNA, ssDNA and dsDNA
     substrates is described. Primer-independent synthesis is
     esp. useful in amplifying RNA for quantitation of RNA species in the
     sample and their identification by direct sequencing. This
     methodol. is esp. useful in detecting pathogenic
     parasites and differences in gene expression levels assocd. with
     diseases.
                                THERE ARE 8 CITED REFERENCES AVAILABLE FOR
REFERENCE COUNT:
                                THIS RECORD. ALL CITATIONS AVAILABLE IN
                                THE RE FORMAT
L16 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER:
                         1997:640790 CAPLUS
DOCUMENT NUMBER:
                         127:289120
                         Nucleotide sequence detection with signal
TITLE:
                         amplification using particle-immobilized probes
                         Delair, Thierry; Elaissari, Abdelhamid; Charles,
INVENTOR(S):
                         Marie-Helene; Mandrand, Bernard
                         Bio Merieux, Fr.; Delair, Thierry; Elaissari,
PATENT ASSIGNEE(S):
                         Abdelhamid; Charles, Marie-Helene; Mandrand,
                         Bernard
SOURCE:
                         PCT Int. Appl., 29 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         French
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                      KIND
                             DATE
                                            APPLICATION NO.
                                            _____
     WO 9735031
                             19970925
                                            WO 1997-FR483
                                                              19970319
                       A1
         W: CA, US
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RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
                                           FR 1996-3412
                                                             19960319
                       A1
                            19970926
     FR 2746413
                            19980424
     FR 2746413
                       В1
                                           US 1997-952397
                                                             19970108
     US 6033853
                            20000307
                       Α
                       AΑ
                            19970925
                                           CA 1997-2219458
                                                            19970319
     CA 2219458
                                           EP 1997-915516
                                                             19970319
     EP 827552
                       A1
                            19980311
                            20020102
     EP 827552
                       В1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
             PT, IE, FI
                                                            19960319
PRIORITY APPLN. INFO.:
                                        FR 1996-3412
                                                         A
                                        WO 1997-FR483
                                                         W 19970319
     A kit for detecting a nucleotide sequence of interest with
AB
     signal amplification is described, said kit contg. a
     labeled nucleotide probe and a reagent including a suspension of
     particles on which at least one series of oligonucleotide units is
     immobilized. Each of the series of oligonucleotide units, which are
     all identical, includes, at least, a nucleotide sequence
     hybridizable with said sequence of interest and a nucleotide
     sequence hybridizable with said probe. Said reagent contains more
     than 10 such oligonucleotide units per particle. The kit
     is particularly useful for diagnosing genetic diseases or
     detg. pathogens such as bacteria, viruses, fungi
     or parasites. A method to detect hepatitis B virus was
     demonstrated. It consisted of a plastic reaction well to which a
     virus-complementary oligonucleotide probe coupled to a polymeric
     tail was passively attached. Viral DNA which hybridized to this
     immobilized probe was detected by addn. of (1) latex particles to
     which many (identical) viral-complementary probes were attached, (2)
     an alk. phosphatase-labeled probe complementary to the latex
     particle-immobilized probes, and (3) a substrate for the
     alk. phosphatase.
L16 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2002 ACS
                         1994:265356 CAPLUS
ACCESSION NUMBER:
                         120:265356
DOCUMENT NUMBER:
                         Detection of Escherichia coli or other
TITLE:
                         microorganism using gas sensors
                         Strachan, Norval James Colin; Ogden, Iain Derek
INVENTOR(S):
                         Minister of Agriculture Fisheries and Food, UK
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 16 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
                         English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                           APPLICATION NO.
                                                             DATE
     PATENT NO.
                      KIND
                            DATE
                                                             19930820
     WO 9404705
                       A1
                            19940303
                                           WO 1993-GB1778
         W: AU, CA, JP, NZ, US
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT,
             SE
                            19950607
                                           EP 1993-919459
                                                             19930820
     EP 656066
                       A1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, NL, PT, SE
                                        GB 1992-17843
                                                             19920821
PRIORITY APPLN. INFO.:
                                        GB 1992-19150
                                                             19920910
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Searcher: Shears 308-4994

WO 1993-GB1778

19930820

AB A method is provided for assessing the contamination status of materials with respect to possible presence of microorganisms (e.g. E. coli), particularly pathogenic microorganisms, by measuring the prodn. of a specific gas or vapor evolved by a sample of the material when it is incubated with a bacterial enzyme substrate. More particularly the method relates to the selective detection of E. coli in the material sample and relating the presence and/or amt. of these to the presence and/or amt. of pathogenic organisms. The method is particularly applicable to test foodstuffs for the likely presence of pathogens. Most preferred substrates are o-nitrophenyl-.beta.-D-glucuronide and methylsalicylyl-.beta.-D-glucuronide. Test kits for carrying out the method are also provided. Foodstuff suspected of contg. contaminating organisms was analyzed. A figure showing the relationship between no. of E. coli colonies found and the incubation time required to obtain a detectable amt. of o-nitrophenol in the head space above an incubated sample is also included.

L16 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1991:625473 CAPLUS

DOCUMENT NUMBER:

115:225473

TITLE:

Procedure for the detection of plant pathogens under field conditions and a diagnostic kit for its application Burochik, Moises; Haim, Liliana

INVENTOR(S): PATENT ASSIGNEE(S): SOURCE:

Agrilab Biotechnology Ltd., Israel

Eur. Pat. Appl., 27 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 444649 EP 444649	A2 A3	19910904 19911227	EP 1991-102956	19910227
R. DE ES	FR. GR	. TT. NI.		

R: DE, ES, FR, GB, IT, NL PRIORITY APPLN. INFO.: AR 1990-316268

A method and a diagnostic kit for detecting phytopathogenic agents in plant tissue by nucleic acid hybridization on a membrane filter under field conditions are described. The method comprises labeling the nucleic acid probes with biotin or sulfonation of the cytosine residues, and detection of the labeling using avidin/streptavidin and the sandwich-type immunoassay, resp., by naked eyes. A set of soln./reagents for hybridization, washing, blocking, and coloring are also disclosed. Detection of potato viruses PVX, PVY, PLRV, and viroid PSTV, and tomato yellow leaf curl virus (TYLCV) were demonstrated.

L16 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2002 ACS 1991:227371 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

114:227371

TITLE:

Monoclonal anti-idiotypic antibodies, their preparation and kits containing them

19900227

for diagnostic detection of antibodies to

infective agents

308-4994 Searcher : Shears